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 Database XReference: WPI; 2000-638353/61.  
 Accession Number: AAH51470  
 Patent Title: Polynucleotides comprising sequences from malate decarboxylase enzyme-related biallelic markers used for genotyping -  
 Patented by: (GEST ) GENSET.  
 Inventor: Blumenfeld M, Bougueleret L, Chumakov I, Cohen-Akenine A  
 Description: Human UGT2B4 related DNA containing a biallelic polymorphism SEQ ID 361.  
 Patent Number: WO200058508-A2  
 Patent Publication Date: 05-OCT-2000  
 Modification Date: 29-AUG-2001 (first entry)  
 Local Filing: 24-MAR-2000; 2000WO-IB00403  
 Priority: 25-MAR-1999  
 Abstract: Sequences AAH51110-AAH51593 represent human DNA fragments which contain biallelic markers. The sequences are related to various human genes including microsomal glutathione S-transferase II (MGSTII), malate decarboxylase enzyme (DME1/ME1), cytochrome P450, glutathione reductase/synthase (GSHR/GSHS), flavin-containing monooxygenases (FMO), gamma-glutamyltransferase 5 (GGT5), dipeptidase (DP), glucose 6-phosphate dehydrogenase (G6PDH), phosphogluconate dehydrogenase (PGDH), and uridine diphosphate glucuronosyl transferases (UGT2). Each of these sequences contains a biallelic marker/polymorphism, which is represented in the sequence as a degenerate/undefined base. The genes to which the biallelic marker containing sequences are related are involved in drug metabolism. Sequences AAH51594 - AAH51598 represent the genomic sequence of the MGSTII gene and four alternative MGSTII cDNA sequences. AAB62905-AAB62906 are MGSTII gene products. PCR primers AAH51599 and AAH51600 are used in an example for the amplification of human genomic DNA fragments. The invention includes a method of genotyping comprising determining the identity of a nucleotide at a DME- or MGSTII-related biallelic marker in a biological sample. The method is used to determine the frequency in population of an allele of a DME- or MGST-II related biallelic marker and to select an individual for inclusion in a clinical trial of a drug treatment. The method is also used to detect association between allele and phenotype, and to detect association between haplotype and phenotype. The polynucleotides are used, in hybridization assays, sequencing assays or allele specific amplification assays. The method can be used to determine whether an individual suffers or is at risk of developing asthma or is at risk of developing hepatotoxicity on treatment with zileuton.

KeyWords: Human;biallelic marker;single nucleotide polymorphism;SNP;MGSTII;microsomal glutathione S-transferase II;malate decarboxylase enzyme;DME1;ME1;cytochrome P450;glutathione reductase;GSHR;GSHS;GGT5;flavin-containing monooxygenase;FMO;gamma-glutamyltransferase 5;dipeptidase;DP;glucose 6-phosphate dehydrogenase;G6PDH;haplotype;phosphogluconate dehydrogenase;PGDH;drug metabolism;phenotype;uridine diphosphate glucuronosyl transferase;UGT2;asthma;hepatotoxicity;zileuton;ds.

Organism: Homo sapiens.

Sequence Composition: Sequence 1001 BP; 331 A; 177 C; 153 G; 300 T; 40 other;

Sequence: >AAH51470 WO200058508-A2 PA (GEST ) PR 25-MAR-1999 PF 24-MAR-2000 Human UGT2B4 related DNA containing a biallelic polymorphism SEQ ID 361. [Homo sapiens.]  
 NNNNTNNNGTGNNNNNNNNNNNCTNNNNANANNNNNNANANNNNCCNNNNGAAGAAAGGC  
 CTGGTGGCCTCTTCTATTCTGGTGCCAGTGCTGCCTCTGAGACACAACAAAGTGATGATG  
 AGAGTTCCTCACATGCAGTTAGAAATAGCACATCAATTTAACAGTGTGATTTTCAGGGCAA  
 TAGGTGTTCCACCTAAACAATNAACCTGAAAGGTACAATTATTCAACAACCTAACTATAAA

CTCTACAATTCCATGTGATAAATGAGACTCCCAAGACTGATTCATAAAAATTCCAAATCA  
CAATACTAGACTCAGGAATGTCAGTGATTCTTAACCACCAGCTTTTATTTTCATTTTTTG  
AAAACTACTGGAAAACCTCTGACAAACTTTAAGTGAAGCATAAAGCATGTAGAGGAACA  
TAAATGTAGATATAAAATTATCCCAACTGTGAATAGCTTTTCCTCAGTGCTCATATTTAG  
GGAAGTAGACCACTAATGKCTTCAAACCTAAAAGAATTCTACAGAAAACCTGCCTGAAATA  
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AAGAACTTATATTAGTAATTATAGTATTATAAGTGAAGAGTCTGGGTATATTTTTTCACA  
TTATCTCCCTGACTACAATGTAATAGCTCCATTTCTTTTCTCCATTACACACATGCAGAC  
ACATACATACATATACACACATATTTACACAAATATCCTTAACAGAGGCCAACTATCTCA  
AATATCTTCTTGCAAAGAACTGAGTGATTGAGTCAGTTAAAAAATATTATTTACTCCAA  
TAATTCCTCAAAATACTTGATTTTCTCTCTTTAATATTTGGTACCAGTTCTTTAGTAGTG  
CCTGCTGTGGTGATACTCTTTTGTGATTAAACAATTTTTTTTTTACAGGAAATGGAGGAG  
TTTGTACAGAGCTCTGGAGAAAATGGTGTTGTGGTGTTTTC



Identifier: AAH51513 DNA Sequence 1001 BP  
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 Patented by: (GEST ) GENSET.  
 Inventor: Blumenfeld M, Bougueleret L, Chumakov I, Cohen-Akenine A  
 Description: Human UGT2B10 related DNA containing a biallelic polymorphism SEQ ID 404.  
 Patent Number: WO200058508-A2  
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 CATTAGTCCAACTGGAAANTCTTGTTATTAANGTTTTGCAGTCTGAAGTCACACCACCATA  
 TAGCCTTCAGTTACATCTCCAACACAAGTACCTGTTTTTNCCTCTGAAATCTGAAAAGT  
 AATAGCAAATTAGTTCAGTGTGTTATCTAGAAAACACTGTCACTTTCAGAGCCTTTTCATT  
 GTGCATCTCATTTTATTCCTATGAATAATTTNTGCTAAAATTCATCCAATCCTAGGTCAT

CCAAAAACCAGAGCTTTTATAACTCATGGTGGAGCCAATGGCATCTATGAGGCAATCTAC  
CATGGGATCCCTATGGTGGGCATTCCATTGTTTTTTNGATCAACCTGATAATATTGCTCA  
CATGAAGGCCAAGGGAGCAGCTGTTAGAGTGGACTTCAACACAATGTCGAGTACAGACCT  
GCTGAATGCACTGAAGACAGTAATTAATGATCCTTCGTGAGTAGAACAATATTTTTCACT  
AGATGGTATTAATAGATAGCTTYTCTTGTCAGTAGTGAGNCATGAGTTTCATCCTTTTTA  
TAAGAGAGTGATTTTGAAAGAATTTAAATGATTTAACCAATCCGAAATCTGCTTTTACTT  
TTTATCTGTTATTTAAAAATTGTATTTGAACCCCATACATCTAATGAGTAACCAGTTAGT  
NGAAACAGTTTTCTAAATAAAAAATAATTTTAAATGATATAGATAATATAAAAAAATACA  
TTTCTTAAAAATTTGACATAATGAATCCATAGTAGAAAGGAAGAATAATCTTGAAATAAT  
ATAATAAAATGTTTTAATTAAATATCTAAATGTCTCAGAATATAACTATTTTCTTGCA  
AAAAATTAATTTTTATTATTATCTTTATTGTAACAGACTTGAAAATGAGATTTAATTTTG  
ATAGCATAAAACCCACCTATTTATGGCAAAAATTCCAAATATTTTACTATGTTTACAGA  
GTCATGAAGTCATCACCAGTGTATAAGTTTGGAACATTTTT



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<b>(21) International Application Number:</b> PCT/IB00/00403 <b>(22) International Filing Date:</b> 24 March 2000 (24.03.00)  <b>(30) Priority Data:</b> 60/126,269      25 March 1999 (25.03.99)      US 60/131,961      30 April 1999 (30.04.99)      US  <b>(71) Applicant (for all designated States except US):</b> GENSET [FR/FR]; Intellectual Property Department, 24, rue Royale, F-75008 Paris (FR).  <b>(72) Inventors; and</b> <b>(75) Inventors/Applicants (for US only):</b> BLUMENFELD, Marta [FR/FR]; 5, rue Tagore, F-75013 Paris (FR). BOUGUEL- ERET, Lydie [FR/FR]; 108, avenue Victor Hugo, F-92170 Vanves (FR). CHUMAKOV, Ilya [FR/FR]; 196, rue des Chevrefoilles, F-77000 Vaux-le-Penil (FR). CO- HEN-AKENINE, Annick [FR/FR]; 76, boulevard Diderot, F-75012 Paris (FR).  <b>(74) Common Representative:</b> GENSET; Intellectual Property De- partment, 24, rue Royale, F-75008 Paris (FR).		<b>(81) Designated States:</b> AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).  <b>Published</b> <i>Without international search report and to be republished          upon receipt of that report.</i>
<b>(54) Title:</b> BIALLELIC MARKERS RELATED TO GENES INVOLVED IN DRUG METABOLISM  <b>(57) Abstract</b>  <p>The invention provides polynucleotides including biallelic markers derived from genes involved in the biotransformation of xenobiotics such as drugs and from genomic regions flanking those genes. Primers hybridizing to regions flanking these biallelic markers are also provided. This invention also provides polynucleotides and methods suitable for genotyping a nucleic acid containing sample for one or more biallelic markers of the invention. Further, the invention provides methods to detect a statistical correlation between a biallelic marker allele and a phenotype and/or between a biallelic marker haplotype and a phenotype.</p>		

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## BIALLELIC MARKERS RELATED TO GENES INVOLVED IN DRUG METABOLISM

### FIELD OF THE INVENTION

5       The present invention is in the field of pharmacogenomics, and is primarily directed to biallelic markers that are located in or in the vicinity of genes, which have an impact on the metabolism of xenobiotics such as drugs and the uses of these markers. The present invention encompasses methods of establishing associations between these markers and a phenotype such as drug response, toxicity and susceptibility to disease. The present  
10 invention also provides means to determine the genetic predisposition of individuals to such drug responses, toxicity and diseases.

### BACKGROUND OF THE INVENTION

To assess the origins of individual variations in drug response, pharmacogenomics uses the genomics technologies to identify polymorphisms within genes associated with drug  
15 response. In this respect, there are three main categories of genes that may theoretically be expected to be associated with drug response, namely genes linked with the targeted disease, genes related to the drug's mode of action and genes involved in the drug's metabolism. Among these genes of pharmacogenomic importance, genes coding for drug-metabolizing enzymes have a central role.

#### 20 **Drug Metabolism**

Drug-metabolizing enzymes are important determinants of drug disposition, safety and efficacy. The enzyme systems involved in the metabolism and the subsequent elimination from the body of environmental chemicals, food toxins and drugs are mainly localized in the liver, although every tissue examined has some metabolic activity.

25       In order to produce its characteristic effects, a given drug must be present in appropriate concentrations at its sites of action. The absorption, distribution, biotransformation and excretion of a drug all involve its passage across cell membranes. The lipophilic characteristics of drugs that promote their passage through biological membranes and subsequent access to their site of action reduce their elimination from the  
30 body. Renal excretion of unchanged drug plays only a modest role in the overall elimination of most therapeutic agents, since lipophilic compounds filtered through the glomerulus are largely reabsorbed through the tubular membranes. Biotransformation of drugs into more hydrophilic metabolites plays a major role in the termination of their biological activity and their elimination from the body. In general, biotransformation

reactions generate more polar, inactive metabolites that are readily excreted from the body. However in some cases, metabolites with potent biological activity or toxic properties are generated and may result in adverse side effects. Metabolic biotransformation of drugs can be classified as either Phase I functionalization reactions or Phase II biosynthetic reactions.

- 5 Phase I reactions introduce or expose a functional group on the parent compound, and generally result in the loss of pharmacological activity although there are some examples of retention or enhancement of activity. Phase II conjugation reactions lead to the formation of a covalent linkage between a functional group on the parent compound with glucuronic acid, sulfate, glutathione, amino acids or acetate. These highly polar conjugates are
- 10 generally inactive and are excreted rapidly in the urine and feces. Within a given cell, most drug metabolizing Phase I enzymes are located primarily in the endoplasmic reticulum, while the Phase II conjugation enzyme systems are mainly cytosolic. In some cases, drugs biotransformed through a Phase I reaction in the endoplasmic reticulum are further metabolized by conjugation in the cytosolic fraction of the same cell (Hardman J.G.,
- 15 Goodman, Gilman A., Limbird L.E.; *Goodman & Gilman's The Pharmacological Basis of Therapeutics*, 9<sup>th</sup> edition, McGraw-Hill, N.Y., 1996).

#### **Enzymes Involved in the Biotransformation of Xenobiotics**

- Besides being involved in the biotransformation of drugs, drug-metabolizing enzymes are also involved in the metabolism of xenobiotics (foreign compounds) as well as
- 20 in the metabolism of endogenous compounds including steroids, vitamins and fatty acids. Foreign compounds include therapeutic agents, carcinogens, plant metabolites, environmental pollutants, foodstuffs and other dietary components as well as industrial chemicals. The biotransformation of foreign compounds (xenobiotics) is often regarded as detoxification because it usually converts compounds into more water-soluble, readily
- 25 excreted substances. This tends to decrease the exposure of the organism to the compound and therefore tends to decrease toxicity. However, in some cases the reverse occurs and a metabolite is produced which is more toxic than the parent compound. For example, drug-metabolizing enzymes may activate some carcinogens, and interindividual differences in cancer susceptibilities, have been linked to polymorphisms in drug-metabolizing enzymes.
- 30 There are many factors, which affect biotransformation and toxicity, such as the dose, availability of cofactors and the relative activity of the various drug-metabolizing enzymes. There may also be several competing pathways of metabolism – some leading to detoxification others to toxicity. Factors, such as genetic factors or environmental factors,

which influence the balance between these competing pathways, will also determine the eventual toxicity.

As mentioned above, the metabolic conversion of drugs and other xenobiotics is enzymatic in nature. The enzyme systems involved in the biotransformation of drugs are localized in the liver, although every tissue examined has some metabolic activity. Other organs with significant metabolic capacity include the kidneys, gastrointestinal tract, skin and lungs. Following non-parenteral administration of a drug, a significant portion of the dose may be metabolically inactivated in either the liver or intestines before it reaches the systemic circulation. This first-pass metabolism significantly limits the oral availability of highly metabolized drugs.

### **Cytochrome P450**

The cytochrome P450 enzyme family is the major catalyst of biotransformation reactions. Since its origin, the cytochrome P450 gene family has diversified to accommodate the metabolism of a growing number of environmental chemicals, food toxins and drugs. The resulting superfamily of enzymes catalyzes a wide variety of oxidative and reductive reactions and has activity towards a chemically diverse group of substrates. Cytochrome P450 enzymes are heme-containing membrane proteins localized in the smooth endoplasmic reticulum of numerous tissues. Oxidative reactions catalyzed by the microsomal monooxygenase system require the cytochrome P450 hemoprotein, NADPH-cytochrome P450 reductase, NADPH, and molecular oxygen. Oxidative biotransformations catalyzed by cytochrome P450 monooxygenases include aromatic and side chain hydroxylation, N-, O-, S- dealkylation, N-oxidation, sulfoxidation, N-hydroxylation, deamination, dehalogenation, and desulfuration. Cytochrome P450 enzymes also catalyze a number of reductive reactions, generally under conditions of low oxygen tension. The only common structural feature of the diverse group of xenobiotics oxidized by cytochrome P450 enzymes is their high lipid solubility.

Twelve cytochrome P450 gene families have been identified in human beings, and a number of distinct cytochrome P450 enzymes often exist within a single cell. The cytochrome P450 1, 2 and 3 families (CYP1, CYP2, CYP3) encode the enzymes involved in the majority of all drug biotransformations, while the gene products of the remaining cytochrome P450 families are important in the metabolism of endogenous compounds such as steroids and fatty acids. CYP1A2 gene expression may play an important role in individual risk of environmental toxicity or cancer. CYP1A2 substrates include clinically important drugs such as imipramine, propranolol, paracetamol, clozapine, theophylline,

caffeine and acetaminophen. CYP1A2 is also involved in the conversion of heterocyclic amines and arylamines to their proximal carcinogenic and mutagenic forms, as well as in the metabolism of endogenous substances including estradiol and uroporphyrinogen III.

Interindividual differences in susceptibility to arylamine- and heterocyclicamine-induced  
5 cancers have been linked to CYP1A2 polymorphism. CYP2C8 appears to be responsible for retinol and retinoic acid metabolism and actively catalyzes benzphetamine N-demethylation. CYP2C9 catalyzes the hydroxylation of tolbutamide, a hypoglycemic agent used in the treatment of type II diabetes mellitus, and one allelic variant of CYP2C9 accounts for the occurrence of poor metabolizers of tolbutamide. CYP2C9 may also have an important role  
10 in terminating the anti-coagulant activity of warfarin. Wide spread interindividual differences in the response to warfarin have been recognized. Such variability is particularly important for drugs such as warfarin which have narrow therapeutic indices (Steward D.J. et al., *Pharmacogenetics*, 7:361-367, 1997). CYP2C9 is further involved in the oxidation of tielinic acid and several non-steroidal anti-inflammatory agents. The  
15 oxidative metabolism through CYP2C9 of tilenic acid can result in the emergence of a drug induced autoimmune hepatitis. CYP3A4 is involved in the biotransformation of a majority of drugs and is expressed at significant levels extrahepatically. It is now recognized that extensive metabolism by CYP3A4 in the gastrointestinal tract is a significant factor contributing to the poor oral availability of many drugs (first-pass metabolism).  
20 Barbiturates, certain steroids and macrolide antibiotics can induce this enzyme. It appears to play a central role in the metabolism of the immunosuppressive cyclic peptide cyclosporin A as well as macrolide antibiotics, such as erythromycin.

#### **Flavin-containing monooxygenases (FMOs)**

The mammalian flavin-containing monooxygenases (FMOs) are microsomal  
25 enzymes that catalyze the NADPH-dependent oxygenation of a wide variety of drugs and other xenobiotics that possess a soft nucleophilic heteroatom, typically a nitrogen, sulfur, phosphorus or selenium atom. Of special clinical interest is the oxidation of trimethylamine in the liver by the FMO, because its deficiency causes the "Fish Odor Syndrome." Drugs oxidized by FMOs include, among others, antidepressant, antipsychotic-neuroleptic,  
30 antihypertensive drugs. FMOs have been implicated in the detoxification but also in the metabolic activation of several different environmental toxins and carcinogens.

Unlike all other known oxidases and monooxygenases, among which the well-studied cytochrome P450 monooxygenases, FMOs have the unique property of forming a stable enzyme intermediate in the absence of an oxygenatable substrate. Because the energy



for catalysis is already present in the FMO enzyme before contact with the potential substrate, the fit of the substrate does not need to be as stringent as with the other enzymes. This feature, unique to FMOs among monooxygenases, is responsible for the wide range of substrates accepted by FMOs (including tertiary and secondary alkyl- and arylamines, many hydrazines, thiocarbamides, thioamides, sulfides, disulfides, thiols, among others), and determines that any soft nucleophilic xenobiotic accessible to the active enzyme will probably be oxidized by FMO *in vivo*. Although some FMO substrates are oxidized to less active derivatives, several soft nucleophiles are metabolized to highly reactive and potentially toxic intermediates.

10       The FMOs represent a multigene family. Five distinct mammalian FMO isoenzymes have been identified and cloned from various animal and human tissues: FMO1, FMO2, FMO3, FMO4 and FMO5. Human FMO2 and human FMOX were cloned and sequenced by the inventors as described in PCT Publication WO 9824914. FMOX represents a new member of the FMO gene family not previously identified in mammals.

15       Tissue specificity and activities of the different FMOs have been thoroughly characterized. FMO1 is known to be expressed in the human kidney but is absent from the liver. In man the enzyme is subject to developmental regulation. FMO2 is predominantly expressed in lung of all mammalian species tested. FMO3 was isolated from human liver, and accounts for the majority of FMO expressed in adult human liver.

20       Many of the FMO substrates may also be oxidized by the cytochrome P450 monooxygenases. However, the final oxidation products are usually different, and the nitrogen of a specific compound is rarely N-oxygenated by both types of monooxygenases. Today, a large number of drugs in human clinical trials contain a nitrogen, sulfur, phosphorous or some other nucleophilic functionality. Of the two major monooxygenase systems considered to be responsible for heteroatom-containing chemical and drug oxidative metabolism (CYP 450 and FMO), relatively little is known concerning the role of the FMO in human drug metabolism. Yet, given the wide range of substrates potentially oxidized by FMOs, this class of monooxygenases seems to represent a major determinant of drug safety and efficacy.

### 30   **Uridine diphosphate glucuronosyl transferase (UGTs)**

Glucuronidation is a major detoxification pathway of Phase II metabolism that is catalyzed by the UDP-glucuronosyl transferase family of enzymes. Glucuronidation is quantitatively the most important conjugation reaction. Members of this enzyme family catalyze the conjugation of numerous endogenous substances of widely differing structures

such as bilirubin, steroid hormones and fat-soluble vitamins. In general, xenobiotics become substrates for glucoronidation by first passing through Phase I metabolism, but many compounds do not require this step because they already possess reactive functionalities (e.g. hydroxyl, carboxyl, amino, sulfhydryl etc.) that are direct targets for glucuronosyl transferase. The human UGT genes appear to have evolved by a series of gene-duplication and gene-conversion events resulting in the emergence of a diversity of isoforms. They are divided into two families, UGT1 which is known to have bilirubin and phenol as substrates, and UGT2 which is known to have steroid, bile, and odorant as substrates, with these two families located on different chromosomes. The UGT2 family is divided into subfamilies UGT2A and UGT2B. The UGTs have different but sometimes overlapping substrate specificities. They catalyze the transfer of an activated glucuronic acid molecule to aromatic and aliphatic alcohols, carboxylic acids, amines and free sulfhydryl groups of both exogenous and endogenous compounds, to form O- N- and S-glucuronide conjugates. The increased water solubility of the glucuronide conjugates promotes their elimination in the urine or bile. In addition to high levels of expression in the liver, UGTs are also found in the kidney, intestine, brain and skin. Glucoronidation constitutes, from a general point of view, a reaction of detoxification and elimination. It generally leads to the formation of inactive metabolites and therefore, glucoronidation can dramatically modify the pharmacological activity of a drug. Moreover, UGTs play a major role in the elimination of nucleophilic metabolites of carcinogens, such as phenols and quinols of polycyclic aromatic hydrocarbons. In this way they prevent their further oxidation to electrophiles, which may react with DNA, RNA or protein. On the other hand, glucoronidation of certain compounds facilitates metabolic activation. Aromatic amines are some of the most studied examples of the role glucoronidation plays in metabolic activation of carcinogens. Glucoronidation has also been implicated in adverse drug reactions of certain carboxylic drugs, which resulted in a toxic immunological response. Glucoronidation although generally a detoxification reaction, may occasionally be involved in increasing toxicity.

#### **Glutathione conjugation and further metabolism**

Glutathione is a tripeptide ( $\gamma$ -glutamylcysteinylglycine, GSH) found in high concentrations in most mammalian tissues, but especially in the liver. Glutathione has several functions including roles in metabolism, transport and catalysis. Glutathione is also important for the maintenance of the thiol moieties of proteins and for the maintenance of the reduced form of other molecules such as cysteine, coenzyme A, and antioxidants such as

ascorbic acid; it is also used in the formation of deoxyribonucleic acids (Anderson M.E., *Advances in Pharmacology*, 38: 65-74, 1997). Glutathione has a major protective role in the body, as it is the major cellular antioxidant. GSH can react non-enzymatically with reactive oxygen species (ROS) and thereby protect the cell from oxidative damage. ROS have been  
5 widely implicated in the pathology of numerous diseases such as arteriosclerosis, rheumatoid arthritis, cancer, AIDS, adult respiratory distress syndrome and Parkinson's disease.

Moreover, conjugation with the tripeptide glutathione represents a major detoxification pathway for xenobiotics including drugs and carcinogens. Glutathione may  
10 react either chemically or in enzyme catalyzed reactions with a variety of compounds, which are reactive electrophilic metabolites produced in Phase I reactions. The glutathione S-transferase enzymes (GSTs) that catalyze these reactions are members of a multigene family and are expressed in virtually all tissues. Glutathione conjugates are cleaved to cysteine derivatives and subsequently are acetylated by a series of enzymes located  
15 primarily in the kidney to give N-acetylcysteine conjugates collectively referred to as mercapturic acids. Mercapturic acid derivatives are the ultimate metabolites excreted in the urine. This is a particularly important route of Phase II metabolism from the toxicological point of view, as it is often involved in the removal of reactive intermediates. Xenobiotics that act as substrates for the glutathione S-transferases (GSTs) fall into four broad  
20 categories: electrophilic carbon, nitrogen, sulfur and oxygen. Examples of substrates for glutathione S-transferases include aromatic, heterocyclic, alicyclic and aliphatic epoxides; aromatic halogen and nitro compounds; alkyl halides; and unsaturated aliphatic compounds (Ballantyne, B., Marrs T. and Turner P., *General & Applied Toxicology*, Stockton Press, New York, 1993). The GSTs are also involved in the metabolism of endogenous molecules  
25 such as the leukotrienes. As mentioned above, many of the enzymes involved in xenobiotic metabolism are also involved in specific aspects of the metabolism of normal cellular biochemical constituents. Leukotrienes are important mediators and modulators of the inflammatory reaction and contribute to a number of physiological and pathological processes. Moreover, the GSTs are also capable of directly binding hydrophobic  
30 compounds such as heme, bilirubin, and steroids, which may enable them to serve as intracellular storage and transport proteins for biological substances with limited water solubility. By their catalytic activity and their capacity for binding, the GSTs provide the cell with mechanism to protect itself from the noxious effects of various xenobiotics and endogenous substances. Further, GSTs may undergo amplification in tumors and may

thereby be implicated in drug resistance in cancer chemotherapy. GSTs are mostly cytosolic although, more recently, microsomal GSTs have been identified. Human microsomal GST II (MGST II) is a member of the microsomal glutathione S-transferase family. This enzyme catalyzes the production of LTC<sub>4</sub> (leukotriene C<sub>4</sub>) from LTA<sub>4</sub> (leukotriene A<sub>4</sub>) and reduced glutathione. Leukotrienes are derived from arachidonic acid and related fatty acids. Metabolites of arachidonic acid have been collectively termed eicosanoids, the principal eicosanoids are prostaglandins, thromboxanes and leukotrienes (LT). Eicosanoids are among the most important chemical mediators and modulators of the inflammatory reaction and contribute to a number of physiological and pathological processes (see Hardman J.G., Goodman, Gilman A., Limbird L.E.; *Goodman & Gilman's The Pharmacological Basis of Therapeutics*, 9<sup>th</sup> edition, McGraw-Hill, N.Y., 1996). The ability to mount an inflammatory response is essential for survival in the face of environmental pathogens and injury, although in some situations and diseases the inflammatory response may be exaggerated and sustained for no apparent beneficial reason. This is the case in numerous chronic inflammatory diseases and allergic inflammation. Acute allergic inflammation is characterized by increased blood flow, extravasation of plasma and recruitment of leukocytes. These events are triggered by locally released inflammatory mediators including eicosanoids and more particularly leukotrienes. The participation of arachidonic acid metabolism in inflammatory diseases such as rheumatoid arthritis, asthma and acute allergy is well established. Pathological actions of leukotrienes are best understood in terms of their roles in immediate hypersensitivity and asthma. LTC<sub>4</sub> and LTD<sub>4</sub> are potent bronchoconstrictors, they act principally on smooth muscle in peripheral airways and are a 1000 times more potent than histamine both *in vitro* and *in vivo*. They also stimulate bronchial mucus secretion and cause mucosal edema. A complex mixture of chemical messengers is released when sensitized lung tissue is challenged by the appropriate antigen. Various prostaglandins and leukotrienes are prominent components of this mixture. Response to the leukotrienes probably dominates during allergic constriction of the airway. A particularly important role for the cysteinyl-leukotrienes (LTC<sub>4</sub>, LTD<sub>4</sub>, and LTE<sub>4</sub>) has been suggested in pathogenesis of asthma, which is now recognized as a chronic inflammatory condition. They are potent spasmogens causing a contraction of bronchiolar muscle and an increase in mucus secretion. An increased LTC<sub>4</sub> formation has also been reported in leukocytes from patients with chronic myelogenous leukemia (Stenke et al., *Acta Oncologica*, 27:803-805, 1987) and in experimental glomerulonephritis (Petric et al., *Biochim. Biophys. Acta*, 1254:207-215, 1995).

Moreover, MGST II has the capacity to conjugate other compounds such as 1-chloro-2, 4 dinitrobenzene with glutathione and may be involved in a general metabolic system for detoxifying fatty acid epoxides (Jakobsson et al., *Journal of Biological Chemistry*, 271:22203-22210, 1996).

5 The resulting glutathione conjugate usually undergoes further metabolism, which involves first a removal of the glutamyl residue, catalyzed by  $\gamma$ -glutamyltransferase (GGT). In addition to catalyzing the initial step in the conversion of glutathione-conjugated compounds to mercapturic acids, GGT also converts LTC<sub>4</sub> to LTD<sub>4</sub>. Interestingly, expression of GGT is often increased in cancerous tissues.

10 Renal dipeptidase is also implicated in the renal metabolism of glutathione and its conjugates including conjugated xenobiotics and endogenous molecules such as Leukotriene D<sub>4</sub>. Pharmacologically it is an important enzyme, for it is responsible for hydrolysis of some  $\beta$ -lactam antibiotics such as penem and carbapenem.

The effectiveness of the GSTs and therefore of detoxification by glutathione  
15 conjugation in general as well as the ability of the cell to resist to oxidative stress, are strongly influenced by the availability of reduced glutathione. Reduction of oxidized glutathione and *de novo* synthesis of glutathione, are both completely dependent on NADPH. Glutathione reductase (GSHR) maintains high levels of reduced glutathione in the cytosol in an NADPH dependent reaction. Reduced glutathione is synthesized *de novo* in  
20 the cytosol of most cells via the  $\gamma$ -glutamyl cycle; a series of tightly controlled, enzyme catalyzed reactions. The first and second step in the *de novo* glutathione biosynthesis are catalyzed by  $\gamma$ -glutamylcysteine synthetase (GLCL) and glutathione synthase (GSHS) respectively. Deficiencies in  $\gamma$ -glutamylcysteine synthetase and GSH synthetase are associated with hemolytic anemia and impaired central nervous system function.

25 **Glucose 6-phosphate dehydrogenase (G6PDH), phosphogluconate dehydrogenase (PGDH) and malate dehydrogenase: generation of NADPH**

NADPH (nicotinamide adenine dinucleotide phosphate) serves as an electron donor in reductive biosyntheses. In the pentose phosphate pathway, NADPH is generated when glucose 6-phosphate is oxidized to ribose 5-phosphate. G6PDH and PGDH are key  
30 enzymes of the pentose phosphate pathway and directly lead to the generation of NADPH. Another major source of NADPH is the oxidative decarboxylation of malate by malic enzyme.

NADPH may be used in anabolic processes such as fatty acid biosynthesis. One of the major functions of malic enzyme may be supplying NADPH to the cytosol for the

synthesis of fatty acids from acetyl CoA (coenzyme A). Further, the cytochrome P450 system is dependent on NADPH. As mentioned above, availability of NADPH is also critical for the reduction of glutathione. The connection between generation of NADPH and reduction of glutathione is clearer in tissues that have limited glycolytic metabolism, e.g. the lens and the erythrocyte. Thus the viability of the erythrocyte depends on glutathione, kept reduced by this pathway. Moreover, factors that influence the availability of reduced glutathione drastically alter the effectiveness of glutathione S-transferases therefore also affecting drug metabolism. Under most conditions saturating levels of NADPH are provided to the cell. However, certain conditions can stress the ability of the cell to provide NADPH and it may become rate limiting.

G6PDH and PGDH are present in most cells and tissues. They serve as the key enzymes of the pentose phosphate pathway that control the flow of carbon through the pathway and produce reducing equivalents as NADPH to meet cellular needs for reductive biosynthesis and to maintain the redox state of the cell at physiological levels. Deficiency of G6PDH and PGDH leads to decreased levels of NADPH and is associated with hemolytic anemia in response to oxidative stress. The red cells of G6PDH deficient persons are susceptible to hemolysis by dietary substances, and by drugs such as primaquine, sulfones, sulfonamides, nitrofurans, vitamin K analogs, acetophenetidin, chloramphenicol, and many others.

## **Genetic Polymorphisms in Drug Metabolizing Enzymes and Pharmacogenomics**

Genetic, environmental, and physiological factors are involved in the regulation of drug biotransformation reactions. Results obtained from epidemiological studies and experimental animal model systems have shown a wide range of phenotypic variation in the ability of individuals to metabolize drugs and environmental chemicals. While some of this variation can be attributed to different environmental exposures, it has become clear that genetic factors also play an important role in determining the response of the individual to exogenous agents. Certain allelic forms of drug-metabolizing enzymes can render the individual either more sensitive or resistant to the toxic or therapeutic effects of exogenous drugs and chemicals. Genetic factors seem to be the major determinants of the variability of drug effects and are responsible for a number of striking quantitative and qualitative differences in pharmacological activity. Genetic differences in the ability of individuals to metabolize a drug through a given pathway are an important contributor to the large interindividual differences of drug efficacy and adverse effects within a population. There are many diverse examples of xenobiotics whose toxicity is directly dependent on the

activity of drug-metabolizing enzymes. Often impaired metabolism of a drug through a genetically polymorphic pathway has been associated with an increased incidence of adverse effects in the slow metabolizer population (Weber W.W., *Pharmacogenetics*, Oxford University Press, N.Y., 1997). Moreover, genetic differences in the regulation, expression and activity of genes coding for Phase I and Phase II drug-metabolizing enzymes can be crucial factors in defining cancer susceptibility and the toxic or carcinogenic power of environmental chemicals and xenobiotics. In addition, the majority of serious cases of drug-drug interactions are a result of the interference of the metabolic clearance of one drug by a coadministered drug. The interference usually occurs via inhibition or induction of drug-metabolizing enzymes. Interindividual differences in susceptibility to severe drug-drug interactions also involve drug-metabolizing enzyme polymorphism. In some cases the design of the drug takes into account the activity of drug-metabolizing enzymes. For example, prodrugs require activation by drug-metabolizing enzymes to exhibit their therapeutic activity. The activation and efficiency of such prodrugs depends on interindividual polymorphism in drug-metabolizing enzymes.

Individual differences in metabolism of therapeutics can lead to severe toxicity or therapeutic failure. Therapeutic management and drug development can be markedly improved by the identification of specific genetic polymorphisms that determine and predict patient susceptibility to diseases or patient responses to drugs. Assessing individual risk rather than population risk will lead to better targeted therapeutic strategies defining individual drug usage based on a benefit/risk prognosis. To assess the origins of individual variations in disease susceptibility or drug response, pharmacogenomics uses the genomic technologies to identify polymorphisms within genes, which are part of biological pathways involved in disease susceptibility, etiology, and development, or more specifically in drug response pathways responsible for a drug's efficacy, tolerance or toxicity. It can provide tools to refine the design of drug development by decreasing the incidence of adverse events in drug tolerance studies, by better defining patient subpopulations of responders and non-responders in efficacy studies and, by combining the results obtained therefrom, to further allow better enlightened individualized drug usage based on efficacy/tolerance prognosis. Pharmacogenomics can also provide tools to identify new targets for designing drugs and to optimize the use of already existing drugs, in order to either increase their response rate and/or exclude non-responders from corresponding treatment, or decrease their undesirable side effects and/or exclude from corresponding treatment patients with marked susceptibility to undesirable side effects.

Drug-metabolizing enzymes are highly relevant to pharmacogenomics because they are at the core of drug response, drug efficacy and toxicity. Drug-metabolizing enzymes also determine an individual's susceptibility to exogenous chemicals and to a number of diseases associated with exposure to toxic or carcinogenic chemicals.

5       The complexity of the pathways and enzymes that are involved in detoxification and metabolism of drugs has limited the precise identification of the drug-metabolizing enzymes, which play the causal role in pathologies or in drug response. Therapeutic management and drug development can be markedly improved by the identification of genetic markers derived from drug-metabolizing enzymes that predict patient susceptibility  
10 to diseases or patient responses to drugs.

#### **Genetic Analysis of Complex Traits**

Until recently, the identification of genes linked with detectable traits has relied mainly on a statistical approach called linkage analysis. Linkage analysis is based upon establishing a correlation between the transmission of genetic markers and that of a specific  
15 trait throughout generations within a family. Linkage analysis involves the study of families with multiple affected individuals and is useful in the detection of inherited traits, which are caused by a single gene, or possibly a very small number of genes. Linkage analysis has been successfully applied to map simple genetic traits that show clear Mendelian inheritance patterns and which have a high penetrance (the probability that a person with a given  
20 genotype will exhibit a trait). About 100 pathological trait-causing genes have been discovered using linkage analysis over the last 10 years.

But, most traits of medical relevance do not follow simple Mendelian monogenic inheritance and linkage studies have proven difficult when applied to complex genetic traits. Many complex traits such as height, blood pressure or cancer susceptibility have been  
25 known to run in families and are at least partially determined by genetic factors. However, the genes or combination of genes that underlie these observable characteristics or traits remain unknown in most cases. Such complex traits are often due to the combined action of multiple genes as well as environmental factors. Because of their low penetrance, such complex traits do not segregate in a clear-cut Mendelian manner as they are passed from one  
30 generation to the next. Drug efficacy, response and tolerance/toxicity can also be considered as multifactoral traits involving a genetic component in the same way as complex diseases. Linkage analysis is impractical when the trait under study is drug response due to the lack of availability of familial cases. In fact, the likelihood of having more than one individual in a family being exposed to the same drug at the same time is very low. Linkage analysis



cannot be applied to the study of such traits for which no large informative families are available. Attempts to map complex traits have been plagued by inconclusive results, demonstrating the need for more sophisticated genetic tools.

Knowledge of genetic variation in drug-metabolizing enzymes is important for understanding why some people are more susceptible to toxicity, pathology or respond differently to drugs. Ways to identify genetic polymorphism and to analyze how they impact and predict disease susceptibility and response to treatment are needed.

Whereas a number of polymorphisms and rare mutations have been identified in drug-metabolizing enzymes (see Weber W.W., *Pharmacogenetics*, Oxford University Press, New York, 1997), genetic markers for use in determining which genes contribute to multigenic or quantitative traits and suitable methods for exploiting those markers have not been found and brought to bare on the genes coding for drug-metabolizing enzymes.

#### **SUMMARY OF THE INVENTION**

The present invention is based on the discovery of a set of novel DME-related biallelic markers. See Figure 1. These markers are located in the coding regions as well as non-coding regions adjacent to genes which are involved in the metabolic conversion of drugs and other xenobiotics. The position of these markers and knowledge of the surrounding sequence has been used to design polynucleotide compositions which are useful in determining the identity of nucleotides at the marker position, as well as more complex association and haplotyping studies which are useful in determining the genetic basis for variability in drug response and adverse reactions to drugs as well as the genetic basis for disease states involving the metabolic conversion of xenobiotics such as drugs. In addition, the compositions and methods of the invention find use in the identification of the targets for the development of pharmaceutical agents and diagnostic methods, as well as the characterization of the differential efficacious responses to and side effects from pharmaceutical agents.

The present invention further stems from the isolation and characterization of the genomic sequence of the MGST-II gene including its regulatory regions and of the complete cDNA sequence encoding the MGST-II enzyme. Oligonucleotide probes and primers hybridizing specifically with a genomic sequence of MGST-II are also part of the invention. A further object of the invention consists of recombinant vectors comprising any of the nucleic acid sequences described in the present invention, and in particular of recombinant vectors comprising the promoter region of MGST-II or a sequence encoding the MGST-II enzyme, as well as cell hosts comprising said nucleic acid sequences or recombinant

vectors. The invention also encompasses methods of screening of molecules which, modulate or inhibit the expression of the MGST-II gene. The invention is also directed to biallelic markers that are located within the MGST-II genomic sequence. these biallelic markers representing useful tools in order to identify a statistically significant association  
5 between specific alleles of MGST-II gene and one or several disorders related to asthma and/or hepatotoxicity.

A first embodiment of the invention encompasses polynucleotides consisting of, consisting essentially of, or comprising a contiguous span of nucleotides of a sequence selected as an individual or in any combination from the group consisting of SEQ ID No. 1-  
10 38, 40-54, 56-463, 465-487, 490-493, the complements thereof, the sequences described in any one or more of Figures 2, 3, 4, 5, 6, 7, and 8, and the complements thereof, wherein said contiguous span is at least 6, 8, 10, 12, 15, 20, 25, 30, 35, 40, 50, 75, 100, 200, 500, or 1000 nucleotides in length, to the extent that such a length is consistent with the lengths of the particular Sequence ID. The present invention also relates to polynucleotides hybridizing  
15 under stringent or intermediate conditions to a sequence selected from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the complements thereof. In addition, the polynucleotides of the invention encompass polynucleotides with any further limitation described in this disclosure, or those following, specified alone or in any combination: Said contiguous span may optionally include the DME-related biallelic marker  
20 in said sequence; Optionally either the original or the alternative allele of Figure 3 may be specified as being present at said DME-related biallelic marker; Optionally either the first or the second allele of Figure 2 or 4 may be specified as being present at said DME-related biallelic marker; Optionally, said polynucleotide may comprise, consists of, or consist essentially of a contiguous span which ranges in length from 8, 10, 12, 15, 18 or 20 to 25,  
25 35, 40, 50, 60, 70, or 80 nucleotides, or be specified as being 12, 15, 18, 20, 25, 35, 40, or 50 nucleotides in length and including a DME-related biallelic marker of said sequence, and optionally the original allele of Figure 3 is present at said biallelic marker; Optionally, said biallelic marker may be within 6, 5, 4, 3, 2, or 1 nucleotides of the center of said polynucleotide or at the center of said polynucleotide; Optionally, the 3' end of said  
30 contiguous span may be present at the 3' end of said polynucleotide; Optionally, biallelic marker may be present at the 3' end of said polynucleotide; Optionally, the 3' end of said polynucleotide may be located within or at least 2, 4, 6, 8, 10, 12, 15, 18, 20, 25, 50, 100, 250, 500, or 1000 nucleotides upstream of a DME-related biallelic marker in said sequence, to the extent that such a distance is consistent with the lengths of the particular Sequence

ID; Optionally, the 3' end of said polynucleotide may be located 1 nucleotide upstream of a DME-related biallelic marker in said sequence; and Optionally, said polynucleotide may further comprise a label.

A second embodiment of the invention encompasses any polynucleotide of the invention attached to a solid support. In addition, the polynucleotides of the invention which are attached to a solid support encompass polynucleotides with any further limitation described in this disclosure, or those following, specified alone or in any combination: Optionally, said polynucleotides may be specified as attached individually or in groups of at least 2, 5, 8, 10, 12, 15, 20, or 25 distinct polynucleotides of the inventions to a single solid support; Optionally, polynucleotides other than those of the invention may be attached to the same solid support as polynucleotides of the invention; Optionally, when multiple polynucleotides are attached to a solid support they may be attached at random locations, or in an ordered array; Optionally, said ordered array may be addressable.

A third embodiment of the invention encompasses the use of any polynucleotide for, or any polynucleotide for use in, determining the identity of one or more nucleotides at a DME-related biallelic marker. In addition, the polynucleotides of the invention for use in determining the identity of one or more nucleotides at a DME-related biallelic marker encompass polynucleotides with any further limitation described in this disclosure, or those following, specified alone or in any combination. Optionally, said DME-related biallelic marker may be in a sequence selected individually or in any combination from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the complements thereof; Optionally, said polynucleotide may comprise a sequence disclosed in the present specification; Optionally, said polynucleotide may consist of, or consist essentially of any polynucleotide described in the present specification; Optionally, said determining may be performed in a hybridization assay, sequencing assay, microsequencing assay, or an enzyme-based mismatch detection assay; Optionally, said polynucleotide may be attached to a solid support, array, or addressable array; Optionally, said polynucleotide may be labeled.

A fourth embodiment of the invention encompasses the use of any polynucleotide for, or any polynucleotide for use in, amplifying a segment of nucleotides comprising a DME-related biallelic marker. In addition, the polynucleotides of the invention for use in amplifying a segment of nucleotides comprising a DME-related biallelic marker encompass polynucleotides with any further limitation described in this disclosure, or those following, specified alone or in any combination: Optionally, said DME-related biallelic marker may

be in a sequence selected individually or in any combination from the group consisting of SEQ ID 1-38, 40-54, 56-463, 465-487, 490-493; and the complements thereof; Optionally, said DME-related biallelic marker may be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker  
5 may be selected from the biallelic markers found in Figures 9, 10, 11 and 12; Optionally, said DME-related biallelic marker may be selected from the following biallelic markers: 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284; Optionally, said  
10 polynucleotide may comprise a sequence disclosed in the present specification; Optionally, said polynucleotide may consist of, or consist essentially of any polynucleotide described in the present specification; Optionally, said amplifying may be performed by a PCR or LCR. Optionally, said polynucleotide may be attached to a solid support, array, or addressable array. Optionally, said polynucleotide may be labeled.

15 A fifth embodiment of the invention encompasses methods of genotyping a biological sample comprising determining the identity of a nucleotide at a DME-related biallelic marker. In addition, the genotyping methods of the invention encompass methods with any further limitation described in this disclosure, or those following, specified alone or in any combination: Optionally, said DME-related biallelic marker may be in a sequence  
20 selected individually or in any combination from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493, and the complements thereof; Optionally, said DME-related biallelic marker may be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may be selected individually or in any combination from the biallelic markers described in  
25 Figure 1; Optionally, said DME-related biallelic marker may be selected from the biallelic markers found in Figures 9, 10, 11 and 12; Optionally, said DME-related biallelic marker may be selected from the following biallelic markers: 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-  
30 219, 12-721-440, and 10-420-284; Optionally, said method further comprises determining the identity of a second nucleotide at said biallelic marker, wherein said first nucleotide and second nucleotide are not base paired (by Watson & Crick base pairing) to one another; Optionally, said biological sample is derived from a single individual or subject; Optionally, said method is performed in vitro; Optionally, said biallelic marker is

determined for both copies of said biallelic marker present in said individual's genome; Optionally, said biological sample is derived from multiple subjects or individuals; Optionally, said method further comprises amplifying a portion of said sequence comprising the biallelic marker prior to said determining step; Optionally, wherein said amplifying is performed by PCR, LCR, or replication of a recombinant vector comprising an origin of replication and said portion in a host cell; Optionally, wherein said determining is performed by a hybridization assay, sequencing assay, microsequencing assay, or an enzyme-based mismatch detection assay.

A sixth embodiment of the invention comprises methods of estimating the frequency of an allele in a population comprising genotyping individuals from said population for a DME-related biallelic marker and determining the proportional representation of said biallelic marker in said population. In addition, the methods of estimating the frequency of an allele in a population of the invention encompass methods with any further limitation described in this disclosure, or those following, specified alone or in any combination:

Optionally, said DME-related biallelic marker may be in a sequence selected individually or in any combination from the group consisting of SEQ No. 1-38, 40-54, 56-463, 465-487, 490-493; and the complements thereof; Optionally, said DME-related biallelic marker may be selected from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may be selected from the biallelic markers found in Figures 9, 10, 11 and 12; Optionally, said DME-related biallelic marker may be selected from the following biallelic markers: 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284; Optionally, determining the frequency of a biallelic marker allele in a population may be accomplished by determining the identity of the nucleotides for both copies of said biallelic marker present in the genome of each individual in said population and calculating the proportional representation of said nucleotide at said DME-related biallelic marker for the population; Optionally, determining the frequency of a biallelic marker allele in a population may be accomplished by performing a genotyping method on a pooled biological sample derived from a representative number of individuals, or each individual, in said population, and calculating the proportional amount of said nucleotide compared with the total.

A seventh embodiment of the invention comprises methods of detecting an association between an allele and a phenotype, comprising the steps of a) determining the frequency of at least one DME-related biallelic marker allele in a case (trait positive) population, b) determining the frequency of said DME-related biallelic marker allele in a control population and; c) determining whether a statistically significant association exists between said genotype and said phenotype. In addition, the methods of detecting an association between an allele and a phenotype of the invention encompass methods with any further limitation described in this disclosure, or those following, specified alone or in any combination: Optionally, said DME-related biallelic marker may be in a sequence selected individually or in any combination from the group consisting of SEQ ID No.1-38, 40-54, 56-463, 465-487, 490-493, and the complements thereof; Optionally, said DME-related biallelic marker may be selected from the biallelic markers described in Figure 1; Optionally, said control population may be a trait negative population, or a random population; Optionally, said phenotype is a response to a drug, or a side effects to a drug, or a disease involving the metabolic conversion of xenobiotics; Optionally, the identity of the nucleotides at the biallelic markers in everyone of the following sequences: SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493 is determined in steps a) and b).

An eighth embodiment of the present invention encompasses methods of estimating the frequency of a haplotype for a set of biallelic markers in a population, comprising the steps of: a) genotyping each individual in said population for at least one DME-related biallelic marker, b) genotyping each individual in said population for a second biallelic marker by determining the identity of the nucleotides at said second biallelic marker for both copies of said second biallelic marker present in the genome; and c) applying a haplotype determination method to the identities of the nucleotides determined in steps a) and b) to obtain an estimate of said frequency. In addition, the methods of estimating the frequency of a haplotype of the invention encompass methods with any further limitation described in this disclosure, or those following, specified alone or in any combination: Optionally said haplotype determination method is selected from the group consisting of asymmetric PCR amplification, double PCR amplification of specific alleles, the Clark method, or an expectation maximization algorithm; Optionally, said second biallelic marker is a DME-related biallelic marker in a sequence selected from the group consisting of the biallelic markers of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493, and the complements thereof; Optionally, said DME-related biallelic markers may be selected individually or in any combination from the biallelic markers described in Figure 1;

Optionally, said DME-related biallelic marker may be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may be selected from the biallelic markers found in Figures 9, 10, 11 and 12; Optionally, said DME-related biallelic marker may be selected from the following

5 biallelic markers: 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284; Optionally, the identity of the nucleotides at the biallelic markers in everyone of the sequences of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493 is determined in steps a)

10 and b).

A ninth embodiment of the present invention encompasses methods of detecting an association between a haplotype and a phenotype, comprising the steps of: a) estimating the frequency of at least one haplotype in a trait positive population according to a method of estimating the frequency of a haplotype of the invention; b) estimating the frequency of said

15 haplotype in a control population according to the method of estimating the frequency of a haplotype of the invention; and c) determining whether a statistically significant association exists between said haplotype and said phenotype. In addition, the methods of detecting an association between a haplotype and a phenotype of the invention encompass methods with any further limitation described in this disclosure, or those following, specified alone or in

20 any combination: Optionally, said DME-related biallelic marker may be in a sequence selected individually or in any combination from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493, and the complements thereof; Optionally, said DME-related biallelic markers may be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may

25 be selected individually or in any combination from the biallelic markers described in Figure 1; Optionally, said DME-related biallelic marker may be selected from the biallelic markers found in Figures 9, 10, 11 and 12; Optionally, said DME-related biallelic marker may be selected from the following biallelic markers: 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-

30 156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284; Optionally, said control population may be a trait negative population, or a random population; Optionally, said phenotype is a response to a drug, or a side effects to a drug, or a disease involving the metabolic conversion of xenobiotics; Optionally, said drug is zileuton and said side effect to said drug is

hepatotoxicity; Optionally, the identity of the nucleotides at the biallelic markers in everyone of the following sequences: SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493 is included in the estimating steps a) and b).

A tenth embodiment of the present invention is a method of administering a drug or  
5 a treatment comprising the steps of: a) obtaining a nucleic acid sample from an individual;  
b) determining the identity of the polymorphic base of at least one DME-related biallelic  
marker which is associated with a positive response to the treatment or the drug; or at least  
one biallelic DME-related biallelic marker which is associated with a negative response to  
the treatment or the drug; and c) administering the treatment or the drug to the individual if  
10 the nucleic acid sample contains said biallelic marker associated with a positive response to  
the treatment or the drug or if the nucleic acid sample lacks said biallelic marker associated  
with a negative response to the treatment or the drug. In addition, the methods of the  
present invention for administering a drug or a treatment encompass methods with any  
further limitation described in this disclosure, or those following, specified alone or in any  
15 combination: optionally, said DME-related biallelic marker may be in a sequence selected  
individually or in any combination from the group consisting of SEQ ID Nos. 1-38, 40-54,  
56-463, 465-487, 490-493; the complements thereof; or preferably SEQ ID Nos. 3, 5, 9, 13-  
15, 25, 31, 33, 37, 38, 41, 323, 345, 351-353, 357, 377, 380; and the complements thereof or  
optionally, the administering step comprises administering the drug or the treatment to the  
20 individual if the nucleic acid sample contains said biallelic marker associated with a positive  
response to the treatment or the drug and the nucleic acid sample lacks said biallelic marker  
associated with a negative response to the treatment or the drug.

An eleventh embodiment of the present invention is a method of selecting an  
individual for inclusion in a clinical trial of a treatment or drug comprising the steps of: a)  
25 obtaining a nucleic acid sample from an individual; b) determining the identity of the  
polymorphic base of at least one DME-related biallelic marker which is associated with a  
positive response to the treatment or the drug, or at least one DME-related biallelic marker  
which is associated with a negative response to the treatment or the drug in the nucleic acid  
sample, and c) including the individual in the clinical trial if the nucleic acid sample  
30 contains said DME-related biallelic marker associated with a positive response to the  
treatment or the drug or if the nucleic acid sample lacks said biallelic marker associated with  
a negative response to the treatment or the drug. In addition, the methods of the present  
invention for selecting an individual for inclusion in a clinical trial of a treatment or drug  
encompass methods with any further limitation described in this disclosure, or those



following, specified alone or in any combination: Optionally, said DME-related biallelic marker may be in a sequence selected individually or in any combination from the group consisting of SEQ ID Nos. 1-38, 40-54, 56-463, 465-487, 490-493; the complements thereof; or preferably SEQ ID Nos. 3, 5, 9, 13-15, 25, 31, 33, 37, 38, 41, 323, 345, 351-353, 357, 377, 380; and the complements thereof, optionally, the including step comprises administering the drug or the treatment to the individual if the nucleic acid sample contains said biallelic marker associated with a positive response to the treatment or the drug and the nucleic acid sample lacks said biallelic marker associated with a negative response to the treatment or the drug.

Additional embodiments are set forth in the Detailed Description of the Invention and in the Examples.

### **BRIEF DESCRIPTION OF THE DRAWINGS**

Figure 1 is a chart containing a list of all of the DME-related biallelic markers for each gene with an indication of the gene for which the marker is in closest physical proximity, an indication of whether the markers have been validated by microsequencing (with a Y indicating that the markers have been validated by microsequencing and an N indicating that it has not), and an indication of the identity and frequency of the least common allele determined by genotyping (with a blank left to indicate that the frequency has not yet been reported for some markers).

Figure 2, 3, and 4 are charts containing lists of the DME-related biallelic markers. Each marker is described by indicating its SEQ ID, the biallelic marker ID, and the two most common alleles. Figure 2 is a chart containing a list of biallelic markers surrounded by preferred sequences. In the column labeled, "POSITION RANGE OF PREFERRED SEQUENCE" of Figure 2, regions of particularly preferred sequences are listed for each SEQ ID, which contain a DME-related biallelic marker, as well as particularly preferred regions of sequences that may not contain a DME-related biallelic marker but, which are in sufficiently close proximity to a DME-related biallelic marker to be useful as amplification or sequencing primers.

Figure 5 is a chart listing particular preferred sequences that are useful for designing some of the primers and probes of the invention. Each sequence is described by indicating its Sequence ID and the positions of the first and last nucleotides (position range) of the particular sequence in the Sequence ID.

Figure 6 is a chart listing microsequencing primers which have been used to genotype DME-related biallelic markers (indicated by an \*) and other preferred

microsequencing primers for use in genotyping DME-related biallelic markers. Each of the primers which falls within the strand of nucleotides included in the Sequence Listing are described by indicating their Sequence ID number and the positions of the first and last nucleotides (position range) of the primers in the Sequence ID. Since the sequences in the Sequence Listing are single stranded and half the possible microsequencing primers are composed of nucleotide sequences from the complementary strand, the primers that are composed of nucleotides in the complementary strand are described by indicating their SEQ ID numbers and the positions of the first and last nucleotides to which they are complementary (complementary position range) in the Sequence ID.

Figure 7 is a chart listing amplification primers which have been used to amplify polynucleotides containing one or more DME-related biallelic markers. Each of the primers which falls within the strand of nucleotides included in the Sequence Listing are described by indicating their Sequence ID number and the positions of the first and last nucleotides (position range) of the primers in the Sequence ID. Since the sequences in the Sequence Listing are single stranded and half the possible amplification primers are composed of nucleotide sequences from the complementary strand, the primers that are composed of nucleotides in the complementary strand are defined by the SEQ ID numbers and the positions of the first and last nucleotides to which they are complementary (complementary position range) in the Sequence ID.

Figure 8 is a chart listing preferred probes useful in genotyping DME-related biallelic markers by hybridization assays. The probes are generally 25-mers with a DME-related biallelic marker in the center position, and described by indicating their Sequence ID number and the positions of the first and last nucleotides (position range) of the probes in the Sequence ID. The probes complementary to the sequences in each position range in each Sequence ID are also understood to be a part of this preferred list even though they are not specified separately.

Figure 9 is a chart containing a list of preferred MGST-II-related biallelic markers with an indication of the frequency of both alleles determined by genotyping. Frequencies were determined in a random US Caucasian population.

Figure 10 is a chart containing a list of preferred ME1-related biallelic markers with an indication of the frequency of both alleles determined by genotyping. Frequencies were determined in a random US Caucasian population.

Figure 11 is a chart containing a list of preferred UGT1A7-related biallelic markers with an indication of the frequency of both alleles determined by genotyping. Frequencies

were determined in a random US Caucasian population and a random French Caucasian population.

Figure 12 is a chart containing a list of preferred UGT2B4-related biallelic markers with an indication of the frequency of both alleles determined by genotyping. Frequencies  
5 were determined in a random US Caucasian population and a random French Caucasian population..

Figure 13 is a chart showing the results of a haplotype analysis study demonstrating an association between asthma and MGST-II-related biallelic marker haplotypes.

Figure 14 is a chart showing the results of a permutation test which evaluates the  
10 statistical significance of the results obtained for the haplotype analysis.

Figure 15 is a chart showing the results of a haplotype analysis study demonstrating an association between side effects upon treatment with the anti-asthmatic drug Zyflo™ (zileuton) and MGST-II related biallelic marker haplotypes.

Figure 16 is a table showing the results of a phenotypic permutation test which  
15 evaluates the statistical significance of the results obtained for the haplotype analysis in Figure 15.

Figure 17 is a block diagram of an exemplary computer system.

Figure 18 is a flow diagram illustrating one embodiment of a process 200 for  
20 comparing a new nucleotide or protein sequence with a database of sequences in order to determine the homology levels between the new sequence and the sequences in the database.

Figure 19 is a flow diagram illustrating one embodiment of a process 250 in a computer for determining whether two sequences are homologous.

Figure 20 is a flow diagram illustrating one embodiment of an identifier process 300  
25 for detecting the presence of a feature in a sequence.

### **BRIEF DESCRIPTION OF THE SEQUENCES PROVIDED IN THE SEQUENCE LISTING**

SEQ ID Nos. 1-484 contain the nucleotide sequence of genomic DNA fragments located in the vicinity of the candidate genes.

30 SEQ ID No. 485 contains the genomic sequence of the MGST-II gene comprising the 5' regulatory region (upstream untranscribed region), the exons and introns, and the 3' regulatory region (downstream untranscribed region).

SEQ ID No. 486 contains a cDNA sequence of MGST-II.

SEQ ID No. 487 contains a cDNA sequence of MGST-II corresponding to an alternative messenger RNA which is due to alternative splicing joining exon 1 to exon 3 and resulting in the absence of exon 2.

SEQ ID No. 488 contains the amino acid sequence encoded by the cDNA of SEQ ID No. 486.

SEQ ID No. 489 contains the amino acid sequence encoded by the cDNA of SEQ ID No. 487.

SEQ ID No. 490 contains a partial cDNA sequence of MGST-II sequence corresponding to a cloned partial messenger RNA.

10 SEQ ID No. 491 contains a partial cDNA sequence of MGST-II sequence corresponding to a cloned partial messenger RNA.

SEQ ID No. 492 contains a primer containing the additional PU 5' sequence described further in Example 1.

SEQ ID No. 493 contains a primer containing the additional RP 5' sequence  
15 described further in Example 1.

### **DETAILED DESCRIPTION OF THE INVENTION**

#### **Advantages of the biallelic markers of the present invention**

The DME-related biallelic markers of the present invention offer a number of important advantages over other genetic markers such as RFLP (Restriction fragment length  
20 polymorphism) and VNTR (Variable Number of Tandem Repeats) markers.

The first generation of markers, were RFLPs, which are variations that modify the length of a restriction fragment. But methods used to identify and to type RFLPs are relatively wasteful of materials, effort, and time. The second generation of genetic markers were VNTRs, which can be categorized as either minisatellites or microsatellites.

25 Minisatellites are tandemly repeated DNA sequences present in units of 5-50 repeats which are distributed along regions of the human chromosomes ranging from 0.1 to 20 kilobases in length. Since they present many possible alleles, their informative content is very high. Minisatellites are scored by performing Southern blots to identify the number of tandem repeats present in a nucleic acid sample from the individual being tested. However, there  
30 are only  $10^4$  potential VNTRs that can be typed by Southern blotting. Moreover, both RFLP and VNTR markers are costly and time-consuming to develop and assay in large numbers.

Single nucleotide polymorphism or biallelic markers can be used in the same manner as RFLPs and VNTRs but offer several advantages. Single nucleotide polymorphisms are densely spaced in the human genome and represent the most frequent type of variation. An

estimated number of more than  $10^7$  sites are scattered along the  $3 \times 10^9$  base pairs of the human genome. Therefore, single nucleotide polymorphism occur at a greater frequency and with greater uniformity than RFLP or VNTR markers which means that there is a greater probability that such a marker will be found in close proximity to a genetic locus of interest. Single nucleotide polymorphisms are less variable than VNTR markers but are mutationally more stable.

Also, the different forms of a characterized single nucleotide polymorphism, such as the biallelic markers of the present invention, are often easier to distinguish and can therefore be typed easily on a routine basis. Biallelic markers have single nucleotide based alleles and they have only two common alleles, which allows highly parallel detection and automated scoring. The biallelic markers of the present invention offer the possibility of rapid, high-throughput genotyping of a large number of individuals.

Biallelic markers are densely spaced in the genome, sufficiently informative and can be assayed in large numbers. The combined effects of these advantages make biallelic markers extremely valuable in genetic studies. Biallelic markers can be used in linkage studies in families, in allele sharing methods, in linkage disequilibrium studies in populations, in association studies of case-control populations. An important aspect of the present invention is that biallelic markers allow association studies to be performed to identify genes involved in complex traits. Association studies examine the frequency of marker alleles in unrelated case- and control-populations and are generally employed in the detection of polygenic or sporadic traits. Association studies may be conducted within the general population and are not limited to studies performed on related individuals in affected families (linkage studies). Biallelic markers in different genes can be screened in parallel for direct association with disease or response to a treatment. This multiple gene approach is a powerful tool for a variety of human genetic studies as it provides the necessary statistical power to examine the synergistic effect of multiple genetic factors on a particular phenotype, drug response, sporadic trait, or disease state with a complex genetic etiology.

#### **Candidate genes of the present invention**

Different approaches can be employed to perform association studies: genome-wide association studies, candidate region association studies and candidate gene association studies. Genome-wide association studies rely on the screening of genetic markers evenly spaced and covering the entire genome. Candidate region association studies rely on the screening of genetic markers evenly spaced covering a region identified as linked to the trait

of interest. The candidate gene approach is based on the study of genetic markers specifically derived from genes potentially involved in a biological pathway related to the trait of interest. In the present invention, genes involved in drug metabolism have been chosen as candidate genes. As mentioned above, these genes are highly relevant to pharmacogenetics because they are at the core of drug response, drug efficacy and toxicity, moreover, drug-metabolizing enzymes also determine an individuals susceptibility to exogenous chemicals and to a number of diseases associated with exposure to toxic or carcinogenic chemicals. The candidate gene analysis clearly provides a short-cut approach to the identification of genes and gene polymorphisms related to a particular trait when some information concerning the biology of the trait is available. However, it should be noted that all of the biallelic markers disclosed in the instant application can be employed as part of genome-wide association studies or as part of candidate region association studies and such uses are specifically contemplated in the present invention and claims. All of the markers are known to be in close proximity to the genes with which they are listed in Figure 1. For a portion of the markers, the precise position of the marker with respect to the various coding and non-coding elements of the genes has also been determined.

The following is a table of abbreviations for the candidate genes as they appear throughout the specification and figures:

**Table 1**

<b>Candidate Gene</b>	<b>Abbreviation</b>
Microsomal glutathione S-transferase II	MGST2 or MGST II
Malate decarboxylase enzyme	DME or ME1
Cytochrome P450 1A2	CYP1A2
Cytochrome P450 2C8	CYP2C8
Cytochrome P450 2C9	CYP2C9
Cytochrome P450 2C18	CYP2C18
Cytochrome P450 3A4-3A7	CYP3A4-CYP3A7
Cytochrome P450 3A7	CYP3A7
Flavin-containing monooxygenases	FMO
Glutathione reductase	GSHR
Glutathione synthase	GSHS
$\gamma$ -glutamylcysteine synthetase	GLCL

$\gamma$ -glutamyltransferase 5	GGT5
Dipeptidase	DP
Glucose 6-phosphate dehydrogenase	G6PDH
Phosphogluconate dehydrogenase	PGDH
Uridine diphosphate glucuronosyl transferase 1A7	UGT1A7
Uridine diphosphate glucuronosyl transferase B4	UGT2B4
Uridine diphosphate glucuronosyl transferase B7	UGT2B7
Uridine diphosphate glucuronosyl transferase B10	UGT2B10
Uridine diphosphate glucuronosyl transferase B15	UGT2B15

### Definitions

As used interchangeably herein, the terms "nucleic acids," "oligonucleotides" and "polynucleotides" include RNA, DNA, or RNA/DNA hybrid sequences of more than one nucleotide in either single chain or duplex form. The term "nucleotide" as used herein as an adjective to describe molecules comprising RNA, DNA, or RNA/DNA hybrid sequences of any length in single-stranded or duplex form. The term "nucleotide" is also used herein as a noun to refer to individual nucleotides or varieties of nucleotides, meaning a molecule, or individual unit in a larger nucleic acid molecule, comprising a purine or pyrimidine, a ribose or deoxyribose sugar moiety, and a phosphate group, or phosphodiester linkage in the case of nucleotides within an oligonucleotide or polynucleotide. Although the term "nucleotide" is also used herein to encompass "modified nucleotides" which comprise at least one modifications (a) an alternative linking group, (b) an analogous form of purine, (c) an analogous form of pyrimidine, or (d) an analogous sugar, for examples of analogous linking groups, purine, pyrimidines, and sugars see for example PCT publication No. WO 95/04064. However, the polynucleotides of the invention are preferably comprised of greater than 50% conventional deoxyribose nucleotides, and most preferably greater than 90% conventional deoxyribose nucleotides. The polynucleotide sequences of the invention may be prepared by any known method, including synthetic, recombinant, *ex vivo* generation, or a combination thereof, as well as utilizing any purification methods known in the art.

Throughout the present specification, the expression "nucleotide sequence" may be employed to designate indifferently a polynucleotide or a nucleic acid. More precisely, the expression "nucleotide sequence" encompasses the nucleic material itself and is thus not

restricted to the sequence information (i.e. the succession of letters chosen among the four base letters) that biochemically characterizes a specific DNA or RNA molecule.

The term "polypeptide" refers to a polymer of amino without regard to the length of the polymer; thus, peptides, oligopeptides, and proteins are included within the definition of polypeptide. This term also does not specify or exclude post-expression modifications of polypeptides, for example, polypeptides which include the covalent attachment of glycosyl groups, acetyl groups, phosphate groups, lipid groups and the like are expressly encompassed by the term polypeptide. Also included within the definition are polypeptides which contain one or more analogs of an amino acid (including, for example, non-naturally occurring amino acids, amino acids which only occur naturally in an unrelated biological system, modified amino acids from mammalian systems etc.), polypeptides with substituted linkages, as well as other modifications known in the art, both naturally occurring and non-naturally occurring.

The term "recombinant polypeptide" is used herein to refer to polypeptides that have been artificially designed and which comprise at least two polypeptide sequences that are not found as contiguous polypeptide sequences in their initial natural environment, or to refer to polypeptides which have been expressed from a recombinant polynucleotide.

The term "isolated" requires that the material be removed from its original environment (e. g., the natural environment if it is naturally occurring). For example, a naturally-occurring polynucleotide or polypeptide present in a living animal is not isolated, but the same polynucleotide or DNA or polypeptide, separated from some or all of the coexisting materials in the natural system, is isolated. Such polynucleotide could be part of a vector and/or such polynucleotide or polypeptide could be part of a composition, and still be isolated in that the vector or composition is not part of its natural environment.

The term "purified" is used herein to describe a polynucleotide or polynucleotide vector of the invention which has been separated from other compounds including, but not limited to other nucleic acids, carbohydrates, lipids and proteins (such as the enzymes used in the synthesis of the polynucleotide), or the separation of covalently closed polynucleotides from linear polynucleotides. A polynucleotide is substantially pure when at least about 50 %, preferably 60 to 75% of a sample exhibits a single polynucleotide sequence and conformation (linear versus covalently close). A substantially pure polynucleotide typically comprises about 50 %, preferably 60 to 90% weight/weight of a nucleic acid sample, more usually about 95%, and preferably is over about 99% pure. Polynucleotide purity or homogeneity may be indicated by a number of means well known



in the art, such as agarose or polyacrylamide gel electrophoresis of a sample, followed by visualizing a single polynucleotide band upon staining the gel. For certain purposes higher resolution can be provided by using HPLC or other means well known in the art.

The term "purified" is further used herein to describe a polypeptide of the invention  
5 which, has been separated from other compounds including, but not limited to nucleic acids, lipids, carbohydrates and other proteins. A polypeptide is substantially pure when at least about 50%, preferably 60 to 75% of a sample exhibits a single polypeptide sequence. A substantially pure polypeptide typically comprises about 50%, preferably 60 to 90% weight/weight of a protein sample, more usually about 95%, and preferably is over about  
10 99% pure. Polypeptide purity or homogeneity is indicated by a number of means well known in the art, such as agarose or polyacrylamide gel electrophoresis of a sample, followed by visualizing a single polypeptide band upon staining the gel. For certain purposes higher resolution can be provided by using HPLC or other means well known in the art.

15 As used herein, the term "non-human animal" refers to any non-human vertebrate, birds and more usually mammals, preferably primates, farm animals such as swine, goats, sheep, donkeys, and horses, rabbits or rodents, more preferably rats or mice. As used herein, the term "animal" is used to refer to any vertebrate, preferable a mammal. Both the terms "animal" and "mammal" expressly embrace human subjects unless preceded with the  
20 term "non-human".

As used herein, the term "antibody" refers to a polypeptide or group of polypeptides which are comprised of at least one binding domain, where an antibody binding domain is formed from the folding of variable domains of an antibody molecule to form three-dimensional binding spaces with an internal surface shape and charge distribution  
25 complementary to the features of an antigenic determinant of an antigen., which allows an immunological reaction with the antigen. Antibodies include recombinant proteins comprising the binding domains, as wells as fragments, including Fab, Fab', F(ab)<sub>2</sub>, and F(ab')<sub>2</sub> fragments.

As used herein, an "antigenic determinant" is the portion of an antigen molecule, in  
30 this case a MGST-II polypeptide, that determines the specificity of the antigen-antibody reaction. An "epitope" refers to an antigenic determinant of a polypeptide. An epitope can comprise as few as 3 amino acids in a spatial conformation which, is unique to the epitope. Generally an epitope consists of at least 6 such amino acids, and more usually at least 8-10 such amino acids. Methods for determining the amino acids which make up an epitope

include x-ray crystallography, 2-dimensional nuclear magnetic resonance, and epitope mapping e.g. the Pepscan method described by H. Mario Geysen et al. 1984. *Proc. Natl. Acad. Sci. U.S.A.* 81:3998-4002; PCT Publication No. WO 84/03564; and PCT Publication No. WO 84/03506.

5       The term "primer" denotes a specific oligonucleotide sequence which is complementary to a target nucleotide sequence and used to hybridize to the target nucleotide sequence. A primer serves as an initiation point for nucleotide polymerization catalyzed by either DNA polymerase, RNA polymerase or reverse transcriptase.

10       The term "probe" denotes a defined nucleic acid segment (or nucleotide analog segment, e.g., polynucleotide as defined herein) which can be used to identify a specific polynucleotide sequence present in samples, said nucleic acid segment comprising a nucleotide sequence complementary of the specific polynucleotide sequence to be identified.

15       The terms "trait" and "phenotype" are used interchangeably herein and refer to any visible, detectable or otherwise measurable property of an organism such as symptoms of, or susceptibility to a disease for example. Typically the terms "trait" or "phenotype" are used herein to refer to symptoms of, or susceptibility to a disease; or to refer to an individual's response to a drug; or to refer to symptoms of, or susceptibility to side effects to a drug. In addition, the terms "trait" or "phenotype" may be used herein to refer to symptoms of, or  
20       susceptibility to a disease involving arachidonic acid metabolism; or to refer to an individual's response to an agent acting on arachidonic acid metabolism; or to refer to symptoms of, or susceptibility to side effects to an agent acting on arachidonic acid metabolism.

25       The term "allele" is used herein to refer to variants of a nucleotide sequence. A biallelic polymorphism has two forms. Typically the first identified allele is designated as the original allele whereas other alleles are designated as alternative alleles. Diploid organisms may be homozygous or heterozygous for an allelic form.

30       The term "heterozygosity rate" is used herein to refer to the incidence of individuals in a population, which are heterozygous at a particular allele. In a biallelic system the heterozygosity rate is on average equal to  $2P_a(1-P_a)$ , where  $P_a$  is the frequency of the least common allele. In order to be useful in genetic studies a genetic marker should have an adequate level of heterozygosity to allow a reasonable probability that a randomly selected person will be heterozygous.

The term "genotype" as used herein refers the identity of the alleles present in an individual or a sample. In the context of the present invention a genotype preferably refers to the description of the biallelic marker alleles present in an individual or a sample. The term "genotyping" a sample or an individual for a biallelic marker consists of determining  
5 the specific allele or the specific nucleotide carried by an individual at a biallelic marker.

The term "mutation" as used herein refers to a difference in DNA sequence between or among different genomes or individuals which has a frequency below 1%.

The term "haplotype" refers to a combination of alleles present in an individual or a sample. In the context of the present invention a haplotype preferably refers to a  
10 combination of biallelic marker alleles found in a given individual and which may be associated with a phenotype.

The term "polymorphism" as used herein refers to the occurrence of two or more alternative genomic sequences or alleles between or among different genomes or individuals. "Polymorphic" refers to the condition in which two or more variants of a  
15 specific genomic sequence can be found in a population. A "polymorphic site" is the locus at which the variation occurs. A single nucleotide polymorphism is a single base pair change. Typically a single nucleotide polymorphism is the replacement of one nucleotide by another nucleotide at the polymorphic site. Deletion of a single nucleotide or insertion of a single nucleotide, also give rise to single nucleotide polymorphisms. In the context of the  
20 present invention "single nucleotide polymorphism" preferably refers to a single nucleotide substitution. Typically, between different genomes or between different individuals, the polymorphic site may be occupied by two different nucleotides.

The terms "biallelic polymorphism" and "biallelic marker" are used interchangeably herein to refer to a polymorphism having two alleles at a fairly high frequency in the  
25 population, preferably a single nucleotide polymorphism. A "biallelic marker allele" refers to the nucleotide variants present at a biallelic marker site. Typically the frequency of the less common allele of the biallelic markers of the present invention has been validated to be greater than 1%, preferably the frequency is greater than 10%, more preferably the frequency is at least 20% (i.e. heterozygosity rate of at least 0.32), even more preferably the  
30 frequency is at least 30% (i.e. heterozygosity rate of at least 0.42). A biallelic marker wherein the frequency of the less common allele is 30% or more is termed a "high quality biallelic marker."

The location of nucleotides in a polynucleotide with respect to the center of the polynucleotide are described herein in the following manner. When a polynucleotide has an

odd number of nucleotides, the nucleotide at an equal distance from the 3' and 5' ends of the polynucleotide is considered to be "at the center" of the polynucleotide, and any nucleotide immediately adjacent to the nucleotide at the center, or the nucleotide at the center itself is considered to be "within 1 nucleotide of the center." With an odd number of nucleotides in  
5 a polynucleotide any of the five nucleotides positions in the middle of the polynucleotide would be considered to be within 2 nucleotides of the center, and so on. When a polynucleotide has an even number of nucleotides, there would be a bond and not a nucleotide at the center of the polynucleotide. Thus, either of the two central nucleotides would be considered to be "within 1 nucleotide of the center" and any of the four  
10 nucleotides in the middle of the polynucleotide would be considered to be "within 2 nucleotides of the center", and so on. For polymorphisms which involve the substitution, insertion or deletion of 1 or more nucleotides, the polymorphism, allele or biallelic marker is "at the center" of a polynucleotide if the difference between the distance from the substituted, inserted, or deleted polynucleotides of the polymorphism and the 3' end of the  
15 polynucleotide, and the distance from the substituted, inserted, or deleted polynucleotides of the polymorphism and the 5' end of the polynucleotide is zero or one nucleotide. If this difference is 0 to 3, then the polymorphism is considered to be "within 1 nucleotide of the center." If the difference is 0 to 5, the polymorphism is considered to be "within 2 nucleotides of the center." If the difference is 0 to 7, the polymorphism is considered to be  
20 "within 3 nucleotides of the center," and so on. For polymorphisms which involve the substitution, insertion or deletion of 1 or more nucleotides, the polymorphism, allele or biallelic marker is "at the center" of a polynucleotide if the difference between the distance from the substituted, inserted, or deleted polynucleotides of the polymorphism and the 3' end of the polynucleotide, and the distance from the substituted, inserted, or deleted  
25 polynucleotides of the polymorphism and the 5' end of the polynucleotide is zero or one nucleotide. If this difference is 0 to 3, then the polymorphism is considered to be "within 1 nucleotide of the center." If the difference is 0 to 5, the polymorphism is considered to be "within 2 nucleotides of the center." If the difference is 0 to 7, the polymorphism is considered to be "within 3 nucleotides of the center," and so on.  
30 The term "upstream" is used herein to refer to a location which, is toward the 5' end of the polynucleotide from a specific reference point.

The terms "base paired" and "Watson & Crick base paired" are used interchangeably herein to refer to nucleotides which can be hydrogen bonded to one another by virtue of their sequence identities in a manner like that found in double-helical DNA with thymine or

uracil residues linked to adenine residues by two hydrogen bonds and cytosine and guanine residues linked by three hydrogen bonds (See Stryer, L., *Biochemistry*, 4th edition, 1995).

The terms "complementary" or "complement thereof" are used herein to refer to the sequences of polynucleotides which is capable of forming Watson & Crick base pairing  
5 with another specified polynucleotide throughout the entirety of the complementary region. This term is applied to pairs of polynucleotides based solely upon their sequences and not any particular set of conditions under which the two polynucleotides would actually bind.

A "promoter" refers to a DNA sequence recognized by the synthetic machinery of the cell required to initiate the specific transcription of a gene.

10 A sequence which is "operably linked" to a regulatory sequence such as a promoter means that said regulatory element is in the correct location and orientation in relation to the nucleic acid to control RNA polymerase initiation and expression of the nucleic acid of interest.

As used herein, the term "operably linked" refers to a linkage of polynucleotide  
15 elements in a functional relationship. For instance, a promoter or enhancer is operably linked to a coding sequence if it affects the transcription of the coding sequence. More precisely, two DNA molecules (such as a polynucleotide containing a promoter region and a polynucleotide encoding a desired polypeptide or polynucleotide) are said to be "operably linked" if the nature of the linkage between the two polynucleotides does not (1) result in  
20 the introduction of a frame-shift mutation or (2) interfere with the ability of the polynucleotide containing the promoter to direct the transcription of the coding polynucleotide.

The terms "disease involving the metabolic conversion of xenobiotics" refers to susceptibility to a condition or to a condition linked to any of the genes listed in Figure 1.  
25 "Disease involving the metabolic conversion of xenobiotics" further refers to a condition involving the biotransformation of drugs and other xenobiotics such as environmental chemicals, food toxins, plant metabolites, carcinogens and industrial chemicals. Such conditions include susceptibility to the toxic or carcinogenic effect of exogenous compounds. "Disease involving the metabolic conversion of xenobiotics" also refers to  
30 disorders in the metabolism of some endogenous compounds such as the metabolism of steroids, vitamins, fatty acids and eicosanoids such as leukotrienes involving any of the drug-metabolizing enzymes shown in Figure 1. "Disease involving the metabolic conversion of xenobiotics" includes, but is not limited to, disorders involving the

cytochrome P450 enzyme family, the flavin containing monooxygenases, glucoronidation, the metabolism of glutathione, the pentose pathway and the generation of NADPH.

The term "disease involving arachidonic acid metabolism" refers to a condition linked to disturbances in expression, production or cellular response to eicosanoids such as  
5 prostaglandins, thromboxanes, prostacyclins, leukotrienes or hydroperoxyeicosatrenoic acids. A disease involving arachidonic acid metabolism further refers to a condition involving one or several enzymes of the distinct enzyme systems contributing to arachidonate metabolism including particularly the 5-lipoxygenase pathway. "Diseases involving arachidonic acid metabolism" also include chronic inflammatory diseases, acute  
10 allergic inflammation and inflammatory conditions such as pain, fever, hypersensitivity, asthma, psoriasis and arthritis. "Diseases involving arachidonic acid metabolism" also include disorders in platelet function, blood pressure, thrombosis, renal function, host defense mechanism, hemostasis, smooth muscle tone, male infertility, primary dysmenorrhea, disorders in parturition, and disorders in tissue injury repair, as well as  
15 disorders in cellular function and development. "Diseases involving arachidonic acid metabolism" also include diseases such as gastrointestinal ulceration, coronary and cerebrovascular syndromes, glomerular immune injury and cancer. Preferably the terms "disease involving arachidonic acid metabolism" refer to a disease including diseases such as cancer, prostate cancer, breast cancer, psoriasis and inflammatory diseases. Preferably  
20 the terms "disease involving arachidonic acid metabolism" refer to a disease involving the 5-lipoxygenase pathway and the biosynthesis of the leukotrienes. More preferably the terms "disease involving arachidonic acid metabolism" refer to a disease involving the synthesis of leukotriene C4 (LTC<sub>4</sub>) and refers to disturbances in expression, activity or function of the human MGST-II enzyme.

25 As used herein the term "DME-related biallelic marker" relates to a set of biallelic markers located in or in the vicinity of the genes disclosed in Figure 1 and further relates to biallelic markers in linkage disequilibrium with the biallelic markers disclosed in Figure 1. The term DME-related biallelic marker encompasses all of the biallelic markers disclosed in Figure 1.

30 The invention also concerns MGST-II-related biallelic markers. The term "MGST-II-related biallelic marker" is used interchangeably herein to relate to all biallelic markers in linkage disequilibrium with the biallelic markers of the MGST-II gene. The term MGST-II-related biallelic marker includes both the genic and non-genic biallelic markers described in Table 2.

The term "non-genic" is used herein to describe MGST-II-related biallelic markers, as well as polynucleotides and primers which occur outside the nucleotide positions shown in the human MGST-II genomic sequence of SEQ ID No. 485. The term "genic" is used herein to describe MGST-II-related biallelic markers as well as polynucleotides and primers  
5 which do occur in the nucleotide positions shown in the human MGST-II genomic sequence of SEQ ID No. 485.

The terms "agent acting on arachidonic acid metabolism" refers to a drug or a compound modulating the activity or concentration of one or several enzymes of the distinct enzyme systems contributing to arachidonate metabolism including particularly the  
10 5-lipoxygenase pathway. "Agent acting on arachidonic acid metabolism" also refers to compounds modulating the formation and action of the eicosanoids including particularly the leukotrienes.

The terms "response to a drug" refer to drug efficacy, including but not limited to ability to metabolize a therapeutic compound, to the ability to convert a pro-drug to an  
15 active drug, and to the pharmacokinetics (absorption, distribution, elimination) and the pharmacodynamics (receptor-related) of a drug in an individual.

The terms "response to an agent acting on arachidonic acid metabolism" refer to drug efficacy, including but not limited to ability to metabolize a compound, to the ability to convert a pro-drug to an active drug, and to the pharmacokinetics (absorption, distribution,  
20 elimination) and the pharmacodynamics (receptor-related) of a drug in an individual.

The terms "side effects to a drug" refer to adverse effects of therapy resulting from extensions of the principal pharmacological action of the drug or to idiosyncratic adverse reactions resulting from an interaction of the drug with unique host factors. "Side effects to a drug" include, but are not limited to, adverse reactions such as dermatologic, hematologic  
25 or hepatologic toxicities and further includes gastric and intestinal ulceration, disturbance in platelet function, renal injury, generalized urticaria, bronchoconstriction, hypotension, and shock.

The terms "side effects to an agent acting on arachidonic acid metabolism" refer to adverse effects of therapy resulting from extensions of the principal pharmacological action  
30 of the drug or to idiosyncratic adverse reactions resulting from an interaction of the drug with unique host factors. The terms "side effects to an agent acting on arachidonic acid metabolism" include, but are not limited to, adverse reactions such as dermatologic, hematologic or hepatologic toxicities.

The term "sequence described in Figure 2" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 2. The SEQ ID that contains each "sequence described in Figure 2" is provided in the column labeled, "SEQ ID NO." The range of nucleotide positions within the Sequence ID of which each sequence consists is provided in the same row as the Sequence ID in a column labeled, "POSITION RANGE OF PREFERRED SEQUENCE". It should be noted that some of the Sequence ID numbers have multiple sequence ranges listed, because they contain multiple "sequences described in Figure 2." Unless otherwise noted the term "sequence described in Figure 2" is to be construed as encompassing sequences that contain either of the two alleles listed in the columns labeled, "1<sup>ST</sup> ALLELE" and "2<sup>ND</sup> ALLELE" at the position identified in field <222> of the allele feature in the appended Sequence Listing for each Sequence ID number referenced in Figure 2.

The term "sequence described in Figure 3" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 3. Unless otherwise noted, the "sequences described in Figure 3" consist of the entire sequence of each Sequence ID provided in the column labeled, "SEQ ID NO." Also unless otherwise noted the term "sequence described in Figure 3" is to be construed as encompassing sequences that contain either of the two alleles listed in the columns labeled, "ORIGINAL ALLELE" and "ALTERNATIVE ALLELE" at the position identified in field <222> of the allele feature in the appended Sequence Listing for each Sequence ID number referenced in Figure 3.

The term "sequence described in Figure 4" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 4. Unless otherwise noted, the "sequences described in Figure 4" consist of the entire sequence of each Sequence ID provided in the column labeled, "SEQ ID NO." Also unless otherwise noted the term "sequence described in Figure 4" is to be construed as encompassing sequences that contain either of the two alleles listed in the columns labeled, "1<sup>ST</sup> ALLELE" and "2<sup>ND</sup> ALLELE" at the position identified in field <222> of the allele feature in the appended Sequence Listing for each Sequence ID number referenced in Figure 4.

The term "sequence described in Figure 5" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 5. The SEQ ID that contains each "sequence described in Figure 5" is provided in the column labeled, "SEQ ID NO." The range of nucleotide positions within the Sequence ID of which each sequence consists is provided in the same row as the Sequence ID in a column labeled,



"POSITION RANGE OF PREFERRED SEQUENCE". It should be noted that some of the Sequence ID numbers have multiple sequence ranges listed, because they contain multiple "sequences described in Figure 5."

The term "sequence described in Figure 6" is used herein to refer to the entire  
 5 collection of nucleotide sequences or any individual sequence defined in Figure 6. The SEQ ID that contains each "sequence described in Figure 6 " is provided in the column labeled, "SEQ ID" The range of nucleotide positions within the Sequence ID of which half of the sequences consists is provided in the same row as the Sequence ID in a column labeled, "POSITION RANGE OF MICROSEQUENCING PRIMERS". The remaining half of the  
 10 sequences described in Figure 6 are complementary to the range of nucleotide positions within the Sequence ID provided in the same row as the Sequence ID in a column labeled, "COMPLEMENTARY POSITION RANGE OF MICROSEQUENCING PRIMERS".

The term "sequence described in Figure 7" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 7. The SEQ  
 15 ID that contains each "sequence described in Figure 7 " is provided in the column labeled, "SEQ ID" The range of nucleotide positions within the Sequence ID of which half of the sequences consists is provided in the same row as the Sequence ID in a column labeled, "POSITION RANGE OF AMPLIFICATION PRIMERS". The remaining half of the sequences described in Figure 7 are complementary to the range of nucleotide positions  
 20 within the Sequence ID provided in the same row as the Sequence ID in a column labeled, "COMPLEMENTARY POSITION RANGE OF AMPLIFICATION PRIMERS".

The term "sequence described in Figure 8" is used herein to refer to the entire collection of nucleotide sequences or any individual sequence defined in Figure 8. The SEQ  
 25 ID that contains each "sequence described in Figure 8 " is provided in the column labeled, "SEQ ID". The range of nucleotide positions within the Sequence ID of which each sequence consists is provided in the same row as the Sequence ID in a column labeled, "POSITION RANGE OF PROBES". The sequences which are complementary to the ranges listed in the column labeled, "POSITION RANGE OF PROBES" are also encompassed by the term, "sequence described in Figure 8." Unless otherwise noted the term "sequence  
 30 described in Figure 8 " is to be construed as encompassing sequences that contain either of the two alleles listed in the allele feature in the sequence listing.

The terms "biallelic marker described in Figure" and "allele described in Figure" are used herein to refer to any or all alleles which are listed in the allele feature in the appended

Sequence Listing for each Sequence ID number referenced in the particular Figure being mentioned.

### **Variants and Fragments**

The invention also relates to variants and fragments of the polynucleotides described  
5 herein, particularly of a MGST-II gene containing one or more biallelic markers according to the invention.

Variants of polynucleotides, as the term is used herein, are polynucleotides that differ from a reference polynucleotide. A variant of a polynucleotide may be a naturally occurring variant such as a naturally occurring allelic variant, or it may be a variant that is  
10 not known to occur naturally. Such non-naturally occurring variants of the polynucleotide may be made by mutagenesis techniques, including those applied to polynucleotides, cells or organisms. Generally, differences are limited so that the nucleotide sequences of the reference and the variant are closely similar overall and, in many regions, identical.

Variants of polynucleotides according to the invention include, without being limited to,  
15 nucleotide sequences which are at least 95% identical, preferably at least 99% identical, more particularly at least 99.5% identical, and most preferably at least 99.8% identical to a polynucleotide selected from the group consisting of the polynucleotides of a sequence from any sequence in the Sequence Listing as well as sequences which are complementary thereto or to any polynucleotide fragment of at least 8 consecutive nucleotides of a sequence  
20 from any sequence in the Sequence Listing. Nucleotide changes present in a variant polynucleotide may be silent, which means that they do not alter the amino acids encoded by the polynucleotide. However, nucleotide changes may also result in amino acid substitutions, additions, deletions, fusions and truncations in the polypeptide encoded by the reference sequence. The substitutions, deletions or additions may involve one or more  
25 nucleotides. The variants may be altered in coding or non-coding regions or both.

Alterations in the coding regions may produce conservative or non-conservative amino acid substitutions, deletions or additions. In the context of the present invention, particularly preferred embodiments are those in which the polynucleotides encode polypeptides which retain substantially the same biological function or activity as the mature MGST-II protein,  
30 or those in which the polynucleotides encode polypeptides which maintain or increase a particular biological activity, while reducing a second biological activity. A polynucleotide fragment is a polynucleotide having a sequence that is entirely the same as part but not all of a given nucleotide sequence, preferably the nucleotide sequence of a MGST-II gene, and variants thereof. The fragment can be a portion of an exon or of an intron of a MGST-II

gene. It can also be a portion of the regulatory regions of MGST-II, preferably of the promoter sequence of the MGST-II gene. Such fragments may be "free-standing", i.e. not part of or fused to other polynucleotides, or they may be comprised within a single larger polynucleotide of which they form a part or region. Indeed, several of these fragments may  
5 be present within a single larger polynucleotide.

#### **Identity Between Nucleic Acids or Polypeptides**

The terms "percentage of sequence identity" and "percentage homology" are used interchangeably herein to refer to comparisons among polynucleotides and polypeptides, and are determined by comparing two optimally aligned sequences over a comparison  
10 window, wherein the portion of the polynucleotide or polypeptide sequence in the comparison window may comprise additions or deletions (i.e., gaps) as compared to the reference sequence (which does not comprise additions or deletions) for optimal alignment of the two sequences. The percentage is calculated by determining the number of positions at which the identical nucleic acid base or amino acid residue occurs in both sequences to  
15 yield the number of matched positions, dividing the number of matched positions by the total number of positions in the window of comparison and multiplying the result by 100 to yield the percentage of sequence identity. Homology is evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA,  
20 and CLUSTALW (Pearson and Lipman, *Proc. Natl. Acad. Sci.* 85(8):2444-2448, 1988; Altschul et al., *J. Mol. Biol.* 215(3):403-410, 1990; Thompson et al., *Nucleic Acids Res.* 22(2):4673-4680, 1994; Higgins et al., *Methods Enzymol.* 266:383-402, 1996; Altschul et al., *Nature Genetics* 3:266-272, 1993). In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search  
25 Tool ("BLAST") which is well known in the art (See, e.g., Karlin and Altschul, *Proc. Natl. Acad. Sci. USA* 87:2267-2268, 1990; Altschul et al., *J. Mol. Biol.* 215(3):403-410, 1990; Altschul et al., *Nature Genetics* 3:266-272, 1993; Altschul et al., *Nuc. Acids Res.* 25:3389-3402, 1997). In particular, five specific BLAST programs are used to perform the following task:

30 (1) BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;  
(2) BLASTN compares a nucleotide query sequence against a nucleotide sequence database;

- (3) BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4) TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
- 5 (5) TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a

10 protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., *Science* 256:1443-1445, 1992; Henikoff and Henikoff, *Proteins* 17:49-61, 1993). Less preferably, the PAM or PAM250 matrices may also be used (See, e.g., Schwartz and Dayhoff, eds.,

15 *Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure*, Washington: National Biomedical Research Foundation, 1978). The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-

20 scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990).

#### **Stringent Hybridization Conditions**

By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h

25 to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 10<sup>6</sup> cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at

30 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2 x SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1 X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15

minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989; and Ausubel et al., 1989. These hybridization conditions are suitable for a nucleic acid molecule of about 20 nucleotides in length. There is no need to say that the hybridization conditions described above are to be adapted according to the length of the desired nucleic acid, following techniques well known to the one skilled in the art. The suitable hybridization conditions may for example be adapted according to the teachings disclosed in the book of Hames and Higgins (*Nucleic Acid Hybridization: A Practical Approach*, IRL Press, Oxford, 1985) or in Sambrook et al. (*Molecular Cloning: A Laboratory Manual*, 2<sup>nd</sup> edition, Cold Spring Harbor Laboratory, Cold Spring Harbor, New York, 1989).

### **I. Biallelic Markers and Polynucleotides Comprising Biallelic Markers**

#### **I.A. Polynucleotides of the Present Invention**

The present invention encompasses polynucleotides for use as primers and probes in the methods of the invention. These polynucleotides may consist of, consist essentially of, or comprise a contiguous span of nucleotides of a sequence from any sequence in the Sequence Listing as well as sequences which are complementary thereto ("complements thereof"). The "contiguous span" may be at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 nucleotides in length, to the extent that a contiguous span of these lengths is consistent with the lengths of the particular Sequence ID. It should be noted that the polynucleotides of the present invention are not limited to having the exact flanking sequences surrounding the polymorphic bases which, are enumerated in the Sequence Listing. Rather, it will be appreciated that the flanking sequences surrounding the biallelic markers, or any of the primers of probes of the invention which, are more distant from the markers, may be lengthened or shortened to any extent compatible with their intended use and the present invention specifically contemplates such sequences. It will be appreciated that the polynucleotides referred to in the Sequence Listing may be of any length compatible with their intended use. Also the flanking regions outside of the contiguous span need not be homologous to native flanking sequences which actually occur in human subjects. The addition of any nucleotide sequence, which is compatible with the nucleotides intended use is specifically contemplated. The contiguous span may optionally include the DME-related biallelic marker in said sequence. Biallelic markers generally consist of a polymorphism at one single base position. Each biallelic marker therefore corresponds to two forms of a polynucleotide sequence which, when compared with one another, present a nucleotide

modification at one position. Usually, the nucleotide modification involves the substitution of one nucleotide for another. Optionally either the original or the alternative allele of the biallelic markers disclosed in Figure 3, or the first or second allele disclosed in Figure 2 and 3 may be specified as being present at the DME-related biallelic marker. Optionally, the biallelic markers may be specified as 12-421-135, 12-442-133, 12-449-63, 12-454-242, 12-463-230, 12-462-199, 10-430-287, 12-718-432, 12-269-301, 2-13-398, 2-28-132, 2-39-27, 2-45-155, 2-4-391, 12-345-410, 10-358-353, 10-360-190, 10-365-374, 10-367-58, 12-468-424, 12-481-293, 12-499-86, 12-500-217, 12-511-101, 12-586-443, 12-593-287, 12-795-383, 10-494-332, 12-659-251, 12-912-419, 12-914-28, 12-624-307 which consist of more complex polymorphisms including insertions/deletions of at least one nucleotide.

Optionally either the original or the alternative allele of these biallelic markers may be specified as being present at the DME-related biallelic marker. Preferred polynucleotides may consist of, consist essentially of, or comprise a contiguous span of nucleotides of a sequence from SEQ ID No. 436-468 well as sequences which are complementary thereto. The "contiguous span" may be at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 nucleotides in length, to the extent that a contiguous span of these lengths is consistent with the lengths of the particular Sequence ID. The contiguous span may optionally comprise a biallelic marker selected from the group consisting of biallelic markers 12-421-135, 12-442-133, 12-449-63, 12-454-242, 12-463-230, 12-462-199, 10-430-287, 12-718-432, 12-269-301, 2-13-398, 2-28-132, 2-39-27, 2-45-155, 2-4-391, 12-345-410, 10-358-353, 10-360-190, 10-365-374, 10-367-58, 12-468-424, 12-481-293, 12-499-86, 12-500-217, 12-511-101, 12-586-443, 12-593-287, 12-795-383, 10-494-332, 12-659-251, 12-912-419, 12-914-28, 12-624-307.

The preferred polynucleotides of the invention include the sequence ranges included in any one the sequence ranges of Figures 2, and 5 to 8 individually or in groups consisting of all the possible combinations of the ranges of included in Figures 2, and 5 to 8. The preferred polynucleotides of the invention also include fragments of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 consecutive nucleotides of the sequence ranges included in any one of the sequence ranges of Figures 2, and 5 to 8 to the extent that fragments of these lengths are consistent with the lengths of the particular sequence range. The preferred polynucleotides of the invention also include fragments of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 consecutive nucleotides of the sequence complementary to the sequence ranges included in any one of the sequence ranges

of Figures 2, and 5 to 8 to the extent that fragments of these lengths are consistent with the lengths of the particular sequence range.

Preferred polynucleotides of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50,  
5 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of a sequence selected from the group consisting of the sequences of SEQ ID Nos. 2-3, 5-30, 437-441, 472, 485-487, and 490-493 and the complements thereof.

Particularly preferred polynucleotides of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30,  
10 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 485, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of the following nucleotide positions of SEQ ID No. 485: 1 to 7667, 7726 to 20264, 20365 to 36918, 36991 to 45180, 45263 to 45741, and 45980 to 49327, and the complements thereof. Other particularly preferred polynucleotides of the present invention include isolated, purified or  
15 recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 486, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of nucleotide positions 1 to 198 of SEQ ID No. 486 and the complements thereof. Other particularly preferred polynucleotides of the present invention include isolated, purified or recombinant  
20 polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 487, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of nucleotide positions 1 to 198 of SEQ ID No. 487 and the complements thereof. Other particularly preferred polynucleotides of the present invention include isolated, purified or recombinant polynucleotides comprising a  
25 contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 490, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of nucleotide positions 1 to 198 of SEQ ID No. 490 and the complements thereof. Other preferred polynucleotides of the present invention include polynucleotides comprising, consisting of, or consisting essentially of a nucleotide sequence  
30 of SEQ ID No. 491.

Particularly preferred polynucleotides of the present invention include purified, isolated or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of a sequence selected from the group consisting of SEQ ID Nos. 3, 5, 9, 13-15, 25, 31, 33, 37,

38, 41, 323, 345, 351-353, 357, 377, and 380, or the complements thereof, wherein said span includes a MGST-II-related biallelic marker. Optionally either allele of the biallelic markers described above in the definition of MGST-II-related biallelic marker is specified as being present at the MGST-II-related biallelic marker.

- 5 Additional preferred polynucleotides of the invention include isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 36971, a C at position 45214 or a T at position 45741 of SEQ ID No. 485. Additional preferred
- 10 polynucleotides of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides from a sequence of SEQ ID No. 486, wherein said contiguous span comprises a T at position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486; or the complement thereof. Additional preferred polynucleotides of the
- 15 invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides from a sequence of SEQ ID No. 487, wherein said contiguous span comprises a T at position 325, a C at position 378 or a T at position 426 of SEQ ID No. 487; or the complements thereof.
- 20 Table 2 contains a list of preferred MGST-II-related biallelic markers. Each marker is described by indicating its Marker ID, the position of the marker in the SEQ ID and the two most common alleles.

**Table 2**

BIALLELIC MARKER ID	ALLELES	POSITION OF BIALLELIC MARKER IN SEQ ID
<b>Non-genic biallelic markers</b>		
12-424-192	A/G	SEQ ID No. 2, position 501
12-424-198	G/T	SEQ ID No. 3, position 501
12-426-154	G/A	SEQ ID No. 5, position 461
12-429-198	C/T	SEQ ID No. 6, position 501
12-430-80	C/T	SEQ ID No. 7, position 501
12-433-215	A/G	SEQ ID No. 8, position 501
12-441-233	A/G	SEQ ID No. 9, position 501
12-441-343	G/A	SEQ ID No. 10, position 501
12-442-221	T/C	SEQ ID No. 11, position 501
12-447-58	G/C	SEQ ID No. 12, position 501
12-449-300	T/C	SEQ ID No. 472, position 501
12-453-429	C/T	SEQ ID No. 13, position 501
12-454-363	A/G	SEQ ID No. 14, position 501



12-455-326	T/C	SEQ ID No. 15, position 501
12-455-383	G/A	SEQ ID No. 16, position 501
12-456-269	A/G	SEQ ID No. 17, position 501
12-456-380	G/T	SEQ ID No. 18, position 501
12-457-204	A/G	SEQ ID No. 19, position 501
12-457-206	C/T	SEQ ID No. 20, position 501
12-458-196	A/T	SEQ ID No. 21, position 501
12-458-438	T/C	SEQ ID No. 22, position 501
12-460-274	A/G	SEQ ID No. 23, position 501
12-461-124	A/C	SEQ ID No. 24, position 501
12-461-299	C/T	SEQ ID No. 25, position 501
12-461-465	C/T	SEQ ID No. 26, position 501
12-462-280	C/T	SEQ ID No. 27, position 501
12-464-66	G/T	SEQ ID No. 28, position 501
12-465-234	G/T	SEQ ID No. 30, position 501
12-465-26	C/T	SEQ ID No. 29, position 501
12-442-133	Deletion G	SEQ ID No. 437, position 501
12-449-63	Insertion AT	SEQ ID No. 438, position 501
10-454-242	Deletion AT	SEQ ID No. 439, position 501
12-463-230	Deletion CAT	SEQ ID No. 440, position 501
12-426-199	Deletion	SEQ ID No. 441, position 501
<b>Genic Biallelic markers</b>		
<b>Biallelic Markers in Genomic sequence (SEQ ID No. 485)</b>		
10-286-289	G/C	7564
10-286-345	A/T	7619
10-286-375	A/G	7649
12-425-57	G/A	17258
12-421-135	insertion of a T	21590
12-421-140	A/G	21595
10-523-232	C/T	36971
10-289-201	C/T	45214
10-290-37	C/T	45741
10-290-326	A/G	46029
10-290-328	G/T	46032
<b>Biallelic Markers in MGST-II cDNA (SEQ ID No. 486)</b>		
10-286-289	G/C	98
10-286-345	A/T	153
10-286-375	A/G	183
10-289-201	C/T	478
10-290-37	C/T	526
<b>Biallelic Markers in MGST-II cDNA (SEQ ID No. 487)</b>		
10-286-289	G/C	98
10-286-345	A/T	153
10-286-375	A/G	183
10-289-201	C/T	378
10-290-37	C/T	426

The invention also relates to polynucleotides that hybridize, under conditions of high or intermediate stringency, to a polynucleotide of a sequence from any sequence in the Sequence Listing as well as sequences, which are complementary thereto. Preferably such

polynucleotides are at least 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 nucleotides in length, to the extent that a polynucleotide of these lengths is consistent with the lengths of the particular Sequence ID. Preferred polynucleotides comprise a DME-related biallelic marker. Optionally either the original or the alternative allele of the biallelic markers

5 disclosed in Figure 3 may be specified as being present at the DME-related biallelic marker. Conditions of high and intermediate stringency are further described in III.C.4 "Methods of Genotyping DNA Samples for Biallelic Markers-Hybridization assay methods."

The present invention further embodies isolated, purified, and recombinant polynucleotides which encode MGST-II polypeptides comprising a contiguous span of at  
10 least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, 30, 40, 50, or 100 amino acids of SEQ ID No. 488. The present invention further embodies isolated, purified, and recombinant polynucleotides which encode the variant MGST-II polypeptides of SEQ ID Nos. 488 and 489. The present invention further embodies isolated, purified, and recombinant polynucleotides which encode a variant  
15 MGST-II polypeptide comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, 30, 40, 50, or 100 amino acids of SEQ ID No. 489. The present invention further embodies isolated, purified, and recombinant polynucleotides which encode polypeptides comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15,  
20 20, 25, 30, 40, 50, or 100 amino acids of SEQ ID No. 488 wherein said contiguous span comprises a His residue at amino acid position 93. The present invention further embodies isolated, purified, and recombinant polynucleotides which encode polypeptides comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, 30, 40, 50, or 100 amino acids of amino acid positions 1-  
25 108 of SEQ ID No. 488.

The primers of the present invention may be designed from the disclosed sequences for any method known in the art. A preferred set of primers is fashioned such that the 3' end of the contiguous span of identity with the sequences of the Sequence Listing is present at the 3' end of the primer. Such a configuration allows the 3' end of the primer to hybridize to  
30 a selected nucleic acid sequence and dramatically increases the efficiency of the primer for amplification or sequencing reactions. In a preferred set of primers the contiguous span is found in one of the sequences described in Figure 5. Allele specific primers may be designed such that a biallelic marker is at the 3' end of the contiguous span and the contiguous span is present at the 3' end of the primer. Such allele specific primers tend to

selectively prime an amplification or sequencing reaction so long as they are used with a nucleic acid sample that contains one of the two alleles present at a biallelic marker. The 3' end of primer of the invention may be located within or at least 2, 4, 6, 8, 10, 12, 15, 18, 20, 25, 50, 100, 250, 500, or 1000, to the extent that this distance is consistent with the particular Sequence ID, nucleotides upstream of a DME-related biallelic marker in said sequence or at any other location which is appropriate for their intended use in sequencing, amplification or the location of novel sequences or markers. A list of preferred amplification primers is disclosed in Figure 7. Primers with their 3' ends located 1 nucleotide upstream of a DME-related biallelic marker have a special utility as microsequencing assays. Preferred microsequencing primers are described in Figure 6.

The probes of the present invention may be designed from the disclosed sequences for any method known in the art, particularly methods which allow for testing if a particular sequence or marker disclosed herein is present. A preferred set of probes may be designed for use in the hybridization assays of the invention in any manner known in the art such that they selectively bind to one allele of a biallelic marker, but not the other under any particular set of assay conditions. Preferred hybridization probes may consists of, consist essentially of, or comprise a contiguous span which ranges in length from 8, 10, 12, 15, 18 or 20 to 25, 35, 40, 50, 60, 70, or 80 nucleotides, or be specified as being 12, 15, 18, 20, 25, 35, 40, or 50 nucleotides in length and including a DME-related biallelic marker of said sequence. Optionally the original allele or alternative allele disclosed in Figure 3 and 4 may be specified as being present at the biallelic marker site. Optionally, said biallelic marker may be within 6, 5, 4, 3, 2, or 1 nucleotides of the center of the hybridization probe or at the center of said probe. A particularly preferred set of hybridization probes is disclosed in Figure 8 or a sequence complementary thereto.

Any of the polynucleotides of the present invention can be labeled, if desired, by incorporating a label detectable by spectroscopic, photochemical, biochemical, immunochemical, or chemical means. For example, useful labels include radioactive substances, fluorescent dyes or biotin. Preferably, polynucleotides are labeled at their 3' and 5' ends. A label can also be used to capture the primer, so as to facilitate the immobilization of either the primer or a primer extension product, such as amplified DNA, on a solid support. A capture label is attached to the primers or probes and can be a specific binding member which forms a binding pair with the solid's phase reagent's specific binding member (e.g. biotin and streptavidin). Therefore depending upon the type of label carried by a polynucleotide or a probe, it may be employed to capture or to detect the target

DNA. Further, it will be understood that the polynucleotides, primers or probes provided herein, may, themselves, serve as the capture label. For example, in the case where a solid phase reagent's binding member is a nucleic acid sequence, it may be selected such that it binds a complementary portion of a primer or probe to thereby immobilize the primer or  
5 probe to the solid phase. In cases where a polynucleotide probe itself serves as the binding member, those skilled in the art will recognize that the probe will contain a sequence or "tail" that is not complementary to the target. In the case where a polynucleotide primer itself serves as the capture label, at least a portion of the primer will be free to hybridize with a nucleic acid on a solid phase. DNA Labeling techniques are well known to the  
10 skilled technician.

Any of the polynucleotides, primers and probes of the present invention can be conveniently immobilized on a solid support. Solid supports are known to those skilled in the art and include the walls of wells of a reaction tray, test tubes, polystyrene beads, magnetic beads, nitrocellulose strips, membranes, microparticles such as latex particles,  
15 sheep (or other animal) red blood cells, duracytes® and others. The solid support is not critical and can be selected by one skilled in the art. Thus, latex particles, microparticles, magnetic or non-magnetic beads, membranes, plastic tubes, walls of microtiter wells, glass or silicon chips, sheep (or other suitable animal's) red blood cells and duracytes are all suitable examples. Suitable methods for immobilizing nucleic acids on solid phases include  
20 ionic, hydrophobic, covalent interactions and the like. A solid support, as used herein, refers to any material which is insoluble, or can be made insoluble by a subsequent reaction. The solid support can be chosen for its intrinsic ability to attract and immobilize the capture reagent. Alternatively, the solid phase can retain an additional receptor which has the ability to attract and immobilize the capture reagent. The additional receptor can include a  
25 charged substance that is oppositely charged with respect to the capture reagent itself or to a charged substance conjugated to the capture reagent. As yet another alternative, the receptor molecule can be any specific binding member which is immobilized upon (attached to) the solid support and which has the ability to immobilize the capture reagent through a specific binding reaction. The receptor molecule enables the indirect binding of the capture  
30 reagent to a solid support material before the performance of the assay or during the performance of the assay. The solid phase thus can be a plastic, derivatized plastic, magnetic or non-magnetic metal, glass or silicon surface of a test tube, microtiter well, sheet, bead, microparticle, chip, sheep (or other suitable animal's) red blood cells, duracytes® and other configurations known to those of ordinary skill in the art. The

polynucleotides of the invention can be attached to or immobilized on a solid support individually or in groups of at least 2, 5, 8, 10, 12, 15, 20, or 25 distinct polynucleotides of the inventions to a single solid support. In addition, polynucleotides other than those of the invention may attached to the same solid support as one or more polynucleotides of the  
5 invention.

Any polynucleotide provided herein may be attached in overlapping areas or at random locations on the solid support. Alternatively the polynucleotides of the invention may be attached in an ordered array wherein each polynucleotide is attached to a distinct region of the solid support which does not overlap with the attachment site of any other  
10 polynucleotide. Preferably, such an ordered array of polynucleotides is designed to be "addressable" where the distinct locations are recorded and can be accessed as part of an assay procedure. Addressable polynucleotide arrays typically comprise a plurality of different oligonucleotide probes that are coupled to a surface of a substrate in different known locations. The knowledge of the precise location of each polynucleotides location  
15 makes these "addressable" arrays particularly useful in hybridization assays. Any addressable array technology known in the art can be employed with the polynucleotides of the invention. One particular embodiment of these polynucleotide arrays is known as the Genechips™, and has been generally described in US Patent 5,143,854; PCT publications WO 90/15070 and 92/10092. These arrays may generally be produced using mechanical  
20 synthesis methods or light directed synthesis methods, which incorporate a combination of photolithographic methods and solid phase oligonucleotide synthesis (Fodor et al., Science, 251:767-777, 1991). The immobilization of arrays of oligonucleotides on solid supports has been rendered possible by the development of a technology generally identified as "Very Large Scale Immobilized Polymer Synthesis" (VLSIPS™) in which, typically, probes are  
25 immobilized in a high density array on a solid surface of a chip. Examples of VLSIPS™ technologies are provided in US Patents 5,143,854 and 5,412,087 and in PCT Publications WO 90/15070, WO 92/10092 and WO 95/11995, which describe methods for forming oligonucleotide arrays through techniques such as light-directed synthesis techniques. In designing strategies aimed at providing arrays of nucleotides immobilized on solid supports,  
30 further presentation strategies were developed to order and display the oligonucleotide arrays on the chips in an attempt to maximize hybridization patterns and sequence information. Examples of such presentation strategies are disclosed in PCT Publications WO 94/12305, WO 94/11530, WO 97/29212 and WO 97/31256.

Oligonucleotide arrays may comprise at least one of the sequences selected from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the sequences complementary thereto or a fragment thereof of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 consecutive nucleotides, to the extent that fragments of these lengths is consistent with the lengths of the particular Sequence ID, for determining whether a sample contains one or more alleles of the biallelic markers of the present invention.

Oligonucleotide arrays may also comprise at least one of the sequences selected from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the sequences complementary thereto or a fragment thereof of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 consecutive nucleotides, to the extent that fragments of these lengths is consistent with the lengths of the particular Sequence ID, for amplifying one or more alleles of the biallelic markers of Figure 1. In other embodiments, arrays may also comprise at least one of the sequences selected from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the sequences complementary thereto or a fragment thereof of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 consecutive nucleotides, to the extent that fragments of these lengths is consistent with the lengths of the particular Sequence ID, for conducting microsequencing analyses to determine whether a sample contains one or more alleles of the biallelic markers of the invention. In still further embodiments, the oligonucleotide array may comprise at least one of the sequences selecting from the group consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493; and the sequences complementary thereto or a fragment thereof of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, 500 or 1000 nucleotides in length, to the extent that fragments of these lengths is consistent with the lengths of the particular Sequence ID, for determining whether a sample contains one or more alleles of the biallelic markers of the present invention.

The present invention also encompasses diagnostic kits comprising one or more polynucleotides of the invention, optionally with a portion or all of the necessary reagents and instructions for genotyping a test subject by determining the identity of a nucleotide at a DME-related biallelic marker. The polynucleotides of a kit may optionally be attached to a solid support, or be part of an array or addressable array of polynucleotides. The kit may provide for the determination of the identity of the nucleotide at a marker position by any method known in the art including, but not limited to, a sequencing assay method, a microsequencing assay method, a hybridization assay method, or an allele specific amplification method. Optionally such a kit may include instructions for scoring the results

of the determination with respect to the test subjects' risk of contracting a diseases involving the metabolic conversion of xenobiotics, or likely response to a drug, or chances of suffering from side effects to a drug, including hepatotoxicity.

#### **I.B. Genomic Sequences of the MGST-II Gene and Biallelic Markers**

5       The present invention encompasses the genomic sequence of the MGST-II gene of SEQ ID No. 485. The MGST-II genomic sequences comprise exons and introns. Particularly preferred genomic sequences of the MGST-II gene of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of  
10   SEQ ID No. 485, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of the following nucleotide positions of SEQ ID No. 485: 1 to 7667, 7726 to 20264, 20365 to 36918, 36991 to 45180, 45263 to 45741, and 45980 to 49327, and the complements thereof. The nucleic acids defining the MGST-II intronic polynucleotides may be used as oligonucleotide primers or probes in order to detect the presence of a copy of the MGST-II  
15   gene in a test sample, or alternatively in order to amplify a target nucleotide sequence within the MGST-II sequences.

The present invention further provides MGST-II intron and exon polynucleotide sequences including biallelic markers. Particularly preferred polynucleotides of the present invention include purified, isolated or recombinant polynucleotides comprising a contiguous  
20   span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of a sequence of SEQ ID No. 485; or the complements thereof; wherein said span includes a MGST-II-related biallelic marker. Preferred polynucleotides comprise at least one biallelic marker selected from the group consisting of biallelic markers 10-286-289, 10-287-116, 10-286-375, 12-425-57, 12-421-135, 12-421-140, 10-523-232, 10-289-201,  
25   10-290-37, 10-290-326 and 10-290-328. The present invention also provides polynucleotides which, may be used as primers and probes in order to amplify fragments carrying biallelic markers or in order to detect biallelic marker alleles.

#### **Regulatory sequences**

The genomic sequence of the MGST-II gene contains regulatory sequences both in  
30   the non-coding 5'- flanking region and in the non-coding 3'- flanking region that border the MGST-II transcribed region containing the 5 exons of this gene. 5'-regulatory sequences of the MGST-II gene comprise the polynucleotide sequences located between the nucleotide in position 1 and the nucleotide in position 7466 of the nucleotide sequence of SEQ ID No. 485. 3'-regulatory sequences of the MGST-II gene comprise the polynucleotide sequences

located between the nucleotide in position 45980 and the nucleotide in position 49327 of the nucleotide sequence of SEQ ID No. 485. Particularly preferred regulatory polynucleotides of the present invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 485, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of the following nucleotide positions of SEQ ID No. 485: 1 to 7466 and 45966 to 49312; and the complements thereof.

The promoter activity of the regulatory regions contained in the MGST-II genomic sequence of polynucleotide sequence of SEQ ID No. 485 can be assessed by any method known in the art. Methods for identifying the polynucleotide fragments of SEQ ID No. 485 involved in the regulation of the expression of the MGST-II gene are well-known to those skilled in the art (see Sambrook et al., Molecular Cloning A Laboratory Manual, Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, 1989). An example of a typical method, that can be used, involves a recombinant vector carrying a reporter gene and genomic sequences from the MGST-II genomic sequence of SEQ ID No. 485. Briefly, the expression of the reporter gene (for example beta galactosidase or chloramphenicol acetyl transferase) is detected when placed under the control of a biologically active polynucleotide fragment. Genomic sequences located upstream of the first exon of the MGST-II gene may be cloned into any suitable promoter reporter vector, such as the pSEAP-Basic, pSEAP-Enhancer, p $\beta$ gal-Basic, p $\beta$ gal-Enhancer, or pEGFP-1 Promoter Reporter vectors available from Clontech, or pGL2-basic or pGL3-basic promoterless luciferase reporter gene vector from Promega. Each of these promoter reporter vectors include multiple cloning sites positioned upstream of a reporter gene encoding a readily assayable protein such as secreted alkaline phosphatase, luciferase, beta galactosidase, or green fluorescent protein. The sequences upstream the first MGST-II exon are inserted into the cloning sites upstream of the reporter gene in both orientations and introduced into an appropriate host cell. The level of reporter protein is assayed and compared to the level obtained with a vector lacking an insert in the cloning site. The presence of an elevated expression level in the vector containing the insert with respect to the control vector indicates the presence of a promoter in the insert.

Promoter sequences within the 5' non-coding regions of the MGST-II gene may be further defined by constructing nested 5' and/or 3' deletions using conventional techniques such as Exonuclease III or appropriate restriction endonuclease digestion. The resulting deletion fragments can be inserted into the promoter reporter vector to determine whether



the deletion has reduced or obliterated promoter activity, such as described, for example, by Coles et al. (*Hum. Mol. Genet.*, 7:791-800, 1998). In this way, the boundaries of the promoters may be defined. If desired, potential individual regulatory sites within the promoter may be identified using site directed mutagenesis or linker scanning to obliterate  
5 potential transcription factor binding sites within the promoter individually or in combination. The effects of these mutations on transcription levels may be determined by inserting the mutations into cloning sites in promoter reporter vectors. This type of assays are well known to those skilled in the art and are further described in WO 97/17359, US 5 374 544, EP 582 796, US 5 698 389, US 5 643 746, US 5 502 176, and US 5 266 488.

10 The activity and the specificity of the promoter of the MGST-II gene can further be assessed by monitoring the expression level of a detectable polynucleotide operably linked to the MGST-II promoter in different types of cells and tissues. The detectable polynucleotide may be either a polynucleotide that specifically hybridizes with a predefined oligonucleotide probe, or a polynucleotide encoding a detectable protein, including a  
15 MGST-II polypeptide or a fragment or a variant thereof. This type of assay is well known to those skilled in the art and is described in US 5 502 176 and US 5 266 488 for example.

Polynucleotides carrying the regulatory elements located both at the 5' end and at the 3' end of the MGST-II coding region may be advantageously used to control the transcriptional and translational activity of an heterologous polynucleotide of interest, said  
20 polynucleotide being heterologous as regards to the MGST-II regulatory region.

Thus, the present invention also concerns a purified, isolated, and recombinant nucleic acid comprising a polynucleotide which, is selected from the group consisting of, the polynucleotide sequences located between the nucleotide in position 1 and the nucleotide in position 7466 of the nucleotide sequence of SEQ ID No. 485; or a sequence  
25 complementary thereto or a biologically active fragment thereof.

By a "biologically active" fragment of SEQ ID No. 485 according to the present invention is intended a polynucleotide comprising or alternatively consisting of a fragment of said polynucleotide which is functional as a regulatory region for expressing a recombinant polypeptide or a recombinant polynucleotide in a recombinant cell host.

30 For the purpose of the invention, a nucleic acid or polynucleotide is "functional" as a regulatory region for expressing a recombinant polypeptide or a recombinant polynucleotide if said regulatory polynucleotide contains nucleotide sequences which contain transcriptional and translational regulatory information, and such sequences are "operably

linked" to nucleotide sequences which encode the desired polypeptide or the desired polynucleotide.

The regulatory polynucleotides according to the invention may be advantageously part of a recombinant expression vector that may be used to express a coding sequence in a  
5 desired host cell or host organism.

#### **I.C. MGST-II cDNA Comprising Biallelic Markers and Variant MGST-II cDNA**

The present invention provides a MGST-II cDNA of SEQ ID No. 486. The Open Reading Frame encoding the MGST-II protein spans from the nucleotide in position 202 to the nucleotide in position 642 of the polynucleotide sequence of SEQ ID No. 486. The  
10 cDNA of SEQ ID No. 486 also includes a 5'-UTR region and a 3'-UTR region.

Particularly preferred cDNA polynucleotides of the present invention include purified, isolated or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of a sequence of SEQ ID No. 486, or the complements thereof, wherein said span includes a  
15 MGST-II-related biallelic marker. Preferred cDNA fragments comprise a biallelic marker selected from the group consisting of 10-286-289 (position 98), 10-286-345 (position 153), 10-286-375 (position 183), 10-523-232 (position 426), 10-289-201 (position 478) and 10-290-37 (position 526). Additional preferred polynucleotides of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least  
20 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides from a sequence of SEQ ID No. 486, wherein said contiguous span comprises a T at position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486; or the complement thereof. Most biallelic polymorphism represent silent nucleotide substitutions but biallelic marker 10-289-201 is associated with an amino acid change in the  
25 corresponding MGST-II polypeptide (TYR replaced by ARG in position 93 of the polypeptide). Moreover, one biallelic marker allele of marker 10-290-37 is associated with a stop codon and the corresponding variant cDNA encodes a truncated MGST-II polypeptide including amino acids 1 to 108.

The present invention further provides a variant MGST-II cDNA of SEQ ID No.  
30 487, corresponding to an alternative splicing form which results in the deletion of exon 2. This alternative splicing of MGST-II yields the variant MGST-II polypeptide of SEQ ID No. 489. MGST-II polypeptides of the present invention are further described below. Preferred cDNAs of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90,

100, 150, 200, 500, or 1000 nucleotides from a sequence of SEQ ID No. 487; or the complements thereof. Additional preferred polynucleotides of the invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides  
5 from a sequence of SEQ ID No. 487, wherein said contiguous span comprises a T at position 325, a C at position 378 or a T at position 426 of SEQ ID No. 487; or the complements thereof. The new exon 1/exon 3 junction sequence of the splice variant MGST-II cDNA, more particularly the nucleotide sequence comprised between the nucleotide in position 106 and the nucleotide in position 374 of the nucleic acid of SEQ ID  
10 No. 486 corresponds to the nucleotide sequence of an EST that has been obtained from a human cDNA library. Polynucleotides comprising this EST of a sequence from SEQ ID No. 490 are also part of the invention.

The above disclosed polynucleotides that contain the coding sequence of the MGST-II gene and of MGST-II variants may be expressed in a desired host cell or a desired host  
15 organism, when this polynucleotide is placed under the control of suitable expression signals. The expression signals may be either the expression signals contained in the regulatory regions in the MGST-II gene of the invention or in contrast the signals may be exogenous regulatory nucleic sequences. Such a polynucleotide, when placed under the suitable expression signals, may also be inserted in a vector for its expression and/or  
20 amplification.

Another preferred cDNA fragment comprises the 5'-UTR region (regulatory) beginning at position 1 and ending at position 201 of SEQ ID Nos. 486 and 487. Preferably said 5'-UTR region comprises a biallelic marker selected from the group consisting of biallelic markers 10-286-289, 10-286-345 and 10-286-375. Particularly preferred 5'-UTR  
25 polynucleotides of the present invention include isolated, purified or recombinant polynucleotides comprising a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500, or 1000 nucleotides of SEQ ID No. 486, wherein said contiguous span comprises at least 1, 2, 3, 4, 5 or 10 of nucleotide positions 1 to 198 of SEQ ID No. 486; and the complements thereof. The 5'-end sequence of the MGST-II cDNA,  
30 more particularly the nucleotide sequence comprised between the nucleotide in position 1 and the nucleotide in position 357 of the nucleic acid of SEQ ID No. 486 corresponds to the nucleotide sequence of a 5'-EST that has been obtained from a human cDNA library. Polynucleotides comprising this 5'-EST of a sequence from SEQ ID No. 490 are also part of the invention.

The polynucleotide disclosed above that contains the coding sequence of the MGST-II gene of the invention may be expressed in a desired host cell or a desired host organism, when this polynucleotide is placed under the control of suitable expression signals. The expression signals may be either the expression signals contained in the regulatory regions 5 in the MGST-II gene of the invention or may be exogenous regulatory nucleic sequences. Such a polynucleotide, when placed under the suitable expression signals, may also be inserted in a vector for its expression.

A further object of the invention consists of an isolated polynucleotide comprising:

- a) a nucleic acid comprising a regulatory nucleotide sequence from a sequence of SEQ ID 10 No. 485;
- b) a polynucleotide encoding a desired polypeptide or a nucleic acid of interest, operably linked to the nucleic acid defined in (a) above;
- c) Optionally, a nucleic acid comprising a 5'-UTR regulatory polynucleotide, preferably a 5'-UTR regulatory polynucleotide sequence of a sequence of SEQ ID No. 486.

15 The polypeptide encoded by the nucleic acid described above may be of various nature or origin, encompassing proteins of prokaryotic or eukaryotic origin. Among the polypeptides expressed under the control of a MGST-II regulatory region, there may be cited bacterial, fungal or viral antigens. Also encompassed are eukaryotic proteins such as intracellular proteins, for example "house keeping" proteins, membrane-bound proteins, for 20 example receptors, and secreted proteins, for example cytokines. In a specific embodiment, the desired polypeptide may be the MGST-II protein, especially the proteins of the amino acid sequence of SEQ ID Nos. 488 and 489.

The desired nucleic acids encoded by the above described polynucleotide, usually a RNA molecule, may be complementary to a desired coding polynucleotide, for example to 25 the MGST-II coding sequence, and thus useful as an antisense polynucleotide.

Such a polynucleotide may be included in a recombinant expression vector in order to express the desired polypeptide or the desired nucleic acid in host cell or in a host organism.

#### **I.D. Polynucleotide Constructs, Recombinant Vectors, Host Cells and Transgenic**

##### **30 Animals**

##### **Polynucleotide Constructs**

The terms "polynucleotide construct" and "recombinant polynucleotide" are used interchangeably herein to refer to linear or circular, purified or isolated polynucleotides that

have been artificially designed and which comprise at least two nucleotide sequences that are not found as contiguous nucleotide sequences in their initial natural environment.

**DNA constructs for expressing the MGST-II gene in recombinant host cells and in transgenic animals**

5 In order to study the physiological and phenotype consequences of a lack of synthesis of the MGST-II protein, both at the cellular level and at the multicellular organism level, in particular as regards to disorders related to abnormal cell proliferation, notably cancers, the invention also encompasses DNA constructs and recombinant vectors enabling a conditional expression of a specific allele of the MGST-II genomic sequence or cDNA

10 A first preferred DNA construct is based on the tetracycline resistance operon *tet* from *E. coli* transposon Tn10 for controlling the MGST-II gene expression, such as described by Gossen et al. (*Science*, 268:1766-1769, 1995). Such a DNA construct contains seven *tet* operator sequences from Tn10 (*tetop*) that are fused to either a minimal promoter or a 5'-regulatory sequence of the MGST-II gene, said minimal promoter or said MGST-II  
15 regulatory sequence being operably linked to a polynucleotide of interest that codes either for a sense or an antisense oligonucleotide or for a polypeptide, including a MGST-II polypeptide or a peptide fragment thereof. This DNA construct is functional as a conditional expression system for the nucleotide sequence of interest when the same cell also comprises a nucleotide sequence coding for either the wild type (tTA) or the mutant (rTA) repressor  
20 fused to the activating domain of viral protein VP16 of herpes simplex virus, placed under the control of a promoter, such as the HCMVIE1 enhancer/promoter or the MMTV-LTR. Indeed, a preferred DNA construct of the invention will comprise both the polynucleotide containing the *tet* operator sequences and the polynucleotide containing a sequence coding for the tTA or the rTA repressor. In the specific embodiment wherein the conditional  
25 expression DNA construct contains the sequence encoding the mutant tetracycline repressor rTA, the expression of the polynucleotide of interest is silent in the absence of tetracycline and induced in its presence.

**DNA constructs allowing homologous recombination: replacement vectors**

A second preferred DNA construct will comprise, from 5'-end to 3'-end : (a) a first  
30 nucleotide sequence that is comprised in the MGST-II genomic sequence; (b) a nucleotide sequence comprising a positive selection marker, such as the marker for neomycine resistance (*neo*); and (c) a second nucleotide sequence that is comprised in the MGST-II genomic sequence, and is located on the genome downstream the first MGST-II nucleotide sequence (a).

In a preferred embodiment, this DNA construct also comprises a negative selection marker located upstream the nucleotide sequence (a) or downstream the nucleotide sequence (c). Preferably, the negative selection marker consists of the thymidine kinase (*tk*) gene (Thomas et al., *Cell*, 44:419-428, 1986), the hygromycin beta gene (Te Riele et al., *Nature*, 348:649-651, 1990), the *hprt* gene (Van der Lugt et al., *Gene*, 105:263-267, 1991; Reid et al., *Proc. Natl. Acad. Sci. USA*, 87:4299-4303, 1990) or the Diphtheria toxin A fragment (*Dt-A*) gene (Nada et al., *Cell*, 73:1125-1135, 1993; Yagi et al., *Proc. Natl. Acad. Sci. USA*, 87:9918-9922, 1990). Preferably, the positive selection marker is located within a MGST-II exon sequence so as to interrupt the sequence encoding a MGST-II protein. These replacement vectors are further described by Mansour et al. (*Nature*, 336:348-352, 1988) and Koller et al. (*Ann. Rev. Immunol.*, 10:705-730, 1992). The first and second nucleotide sequences (a) and (c) may be indifferently located within a MGST-II regulatory sequence, an intronic sequence, an exon sequence or a sequence containing both regulatory and/or intronic and/or exon sequences. The size of the nucleotide sequences (a) and (c) is ranging from 1 to 50 kb, preferably from 1 to 10 kb, more preferably from 2 to 6 kb and most preferably from 2 to 4 kb.

#### **DNA constructs allowing homologous recombination: Cre-loxP system**

These new DNA constructs make use of the site specific recombination system of the P1 phage. The P1 phage possesses a recombinase called Cre which, interacts specifically with a 34 base pairs loxP site. The loxP site is composed of two palindromic sequences of 13 bp separated by a 8 bp conserved sequence (Hoess et al., *Nucleic Acids Res.*, 14:2287-2300, 1986). The recombination by the Cre enzyme between two loxP sites having an identical orientation leads to the deletion of the DNA fragment.

The Cre-loxP system used in combination with a homologous recombination technique has been first described by Gu et al. (*Cell*, 73:1155-1164, 1993). Briefly, a nucleotide sequence of interest to be inserted in a targeted location of the genome harbors at least two loxP sites in the same orientation and located at the respective ends of a nucleotide sequence to be excised from the recombinant genome. The excision event requires the presence of the recombinase (Cre) enzyme within the nucleus of the recombinant cell host. The recombinase enzyme may be brought at the desired time either by (a) incubating the recombinant cell hosts in a culture medium containing this enzyme, by injecting the Cre enzyme directly into the desired cell, such as described by Araki et al. (*Proc. Natl. Acad. Sci. USA*, 92: 160-164, 1995), or by lipofection of the enzyme into the cells, such as described by Baubonis et al. (*Nucleic Acids Res.*, 21:2025-2029, 1993); (b) transfecting the

- cell host with a vector comprising the *Cre* coding sequence operably linked to a promoter functional in the recombinant cell host, which promoter being optionally inducible, said vector being introduced in the recombinant cell host, such as described by Gu et al. (*Cell*, 73:1155-1164, 1993) and Sauer et al. (*Proc. Natl. Acad. Sci. USA*, 85:5166-5170, 1988); (c) introducing in the genome of the cell host a polynucleotide comprising the *Cre* coding sequence operably linked to a promoter functional in the recombinant cell host, which promoter is optionally inducible, and said polynucleotide being inserted in the genome of the cell host either by a random insertion event or an homologous recombination event, such as described by Gu et al. (*Science*, 265:103-106, 1994).
- 10 In the specific embodiment wherein the vector containing the sequence to be inserted in the MGST-II gene by homologous recombination is constructed in such a way that selectable markers are flanked by *loxP* sites of the same orientation, it is possible, by treatment by the Cre enzyme, to eliminate the selectable markers while leaving the MGST-II sequences of interest that have been inserted by an homologous recombination event.
- 15 Again, two selectable markers are needed: a positive selection marker to select for the recombination event and a negative selection marker to select for the homologous recombination event. Vectors and methods using the Cre-*loxP* system are further described by Zou et al. (*Curr. Biol.*, 4:1099-1103, 1994).

- Thus, a third preferred DNA construct of the invention comprises, from 5'-end to 3'-end: (a) a first nucleotide sequence that is comprised in the MGST-II genomic sequence; (b) a nucleotide sequence comprising a polynucleotide encoding a positive selection marker, said nucleotide sequence comprising additionally two sequences defining a site recognized by a recombinase, such as a *loxP* site, the two sites being placed in the same orientation; and (c) a second nucleotide sequence that is comprised in the MGST-II genomic sequence, and
- 20 is located on the genome downstream of the first MGST-II nucleotide sequence (a).

- The sequences defining a site recognized by a recombinase, such as a *loxP* site, are preferably located within the nucleotide sequence (b) at suitable locations bordering the nucleotide sequence for which the conditional excision is sought. In one specific embodiment, two *loxP* sites are located at each side of the positive selection marker
- 30 sequence, in order to allow its excision at a desired time after the occurrence of the homologous recombination event.

In a preferred embodiment of a method using the third DNA construct described above, the excision of the polynucleotide fragment bordered by the two sites recognized by a recombinase, preferably two *loxP* sites, is performed at a desired time. due to the presence

within the genome of the recombinant cell host of a sequence encoding the Cre enzyme operably linked to a promoter sequence, preferably an inducible promoter, more preferably a tissue-specific promoter sequence and most preferably a promoter sequence which is both inducible and tissue-specific, such as described by Gu et al. (*Science*, 265:103-106, 1994).

5 The presence of the Cre enzyme within the genome of the recombinant cell host may result of the breeding of two transgenic animals, the first transgenic animal bearing the MGST-II-derived sequence of interest containing the *loxP* sites as described above and the second transgenic animal bearing the *Cre* coding sequence operably linked to a suitable promoter sequence, such as described by Gu et al. (*Science*, 265:103-106, 1994).

10 Spatio-temporal control of the Cre enzyme expression may also be achieved with an adenovirus based vector that contains the Cre gene thus allowing infection of cells, or *in vivo* infection of organs, for delivery of the Cre enzyme, such as described by Anton et al. (*J. Virol.*, 69:4600-4606, 1995) and Kanegae et al. (*Nucleic Acids Res.*, 23:3816-3821, 1995).

15 The DNA constructs described above may be used to introduce a desired nucleotide sequence of the invention, preferably a MGST-II genomic sequence or a MGST-II cDNA sequence, and most preferably an altered copy of a MGST-II genomic or cDNA sequence, within a predetermined location of the targeted genome, leading either to the generation of an altered copy of a targeted gene (knock-out homologous recombination) or to the  
20 replacement of a copy of the targeted gene by another copy sufficiently homologous to allow an homologous recombination event to occur (knock-in homologous recombination).

#### **Recombinant Vectors**

The term "vector" is used herein to designate either a circular or a linear DNA or RNA molecule, which is either double-stranded or single-stranded, and which comprise at  
25 least one polynucleotide of interest that is sought to be transferred in a cell host or in a unicellular or multicellular host organism.

The present invention encompasses a family of recombinant vectors that comprise a regulatory polynucleotide derived from the MGST-II genomic sequence, or a coding polynucleotide from the MGST-II genomic sequence. Consequently, the present invention  
30 further deals with a recombinant vector comprising either a regulatory polynucleotide comprised in the nucleic acid of SEQ ID Nos. 485 and 486 or a polynucleotide comprising the MGST-II coding sequence or both.

In a first preferred embodiment, a recombinant vector of the invention is used to amplify the inserted polynucleotide derived from a MGST-II genomic sequence selected



from the group consisting of the nucleic acids of SEQ ID No. 485 or a MGST-II cDNA, for example the cDNA of SEQ ID Nos. 486 and 487 in a suitable host cell, this polynucleotide being amplified each time the recombinant vector replicates. Generally, a recombinant vector of the invention may comprise any of the polynucleotides described herein, including  
5 regulatory sequences and coding sequences, as well as any MGST-II primer or probe as defined above.

A second preferred embodiment of the recombinant vectors according to the invention consists of expression vectors comprising either a regulatory polynucleotide or a coding nucleic acid of the invention, or both. Within certain embodiments, expression  
10 vectors are employed to express the MGST-II polypeptide which can be then purified and, for example be used in ligand screening assays or as an immunogen in order to raise specific antibodies directed against the MGST-II protein. In other embodiments, the expression vectors are used for constructing transgenic animals and also for gene therapy. Expression requires that appropriate signals are provided in the vectors, said signals including various  
15 regulatory elements, such as enhancers/promoters from both viral and mammalian sources that drive expression of the genes of interest in host cells. Dominant drug selection markers for establishing permanent, stable cell clones expressing the products are generally included in the expression vectors of the invention, as they are elements that link expression of the drug selection markers to expression of the polypeptide.

20 More particularly, the present invention relates to expression vectors which include nucleic acids encoding a MGST-II protein, preferably the MGST-II protein of the amino acid sequence of SEQ ID Nos. 488 and 489, under the control of a regulatory sequence selected among the MGST-II regulatory polynucleotides of SEQ ID Nos. 485 and 486, or alternatively under the control of an exogenous regulatory sequence. Consequently,  
25 preferred expression vectors of the invention are selected from the group consisting of : (a) the MGST-II regulatory sequence comprised therein drives the expression of a coding polynucleotide operably linked thereto; (b) the MGST-II coding sequence is operably linked to regulation sequences allowing its expression in a suitable cell host and/or host organism. Additionally, the recombinant expression vector described above may also comprise a  
30 nucleic acid comprising a 5'-regulatory polynucleotide or a 3'-regulatory polynucleotide, preferably a 5'-regulatory polynucleotide or a 3'-regulatory polynucleotide of the MGST-II gene. The MGST-II 5'-regulatory polynucleotide may also comprise the 5'-UTR sequence contained in the nucleotide sequence of SEQ ID Nos. 486 and 487; or a biologically active fragment or variant thereof. The invention also pertains to a recombinant expression vector

useful for the expression of the MGST-II coding sequence, wherein said vector comprises any of the MGST-II cDNAs or cDNA variants described above; or fragments thereof.

Some of the elements which, can be found in the vectors of the present invention are described in further detail in the following sections.

5 1. General features of the expression vectors of the invention

- A recombinant vector according to the invention comprises, but is not limited to, a YAC (Yeast Artificial Chromosome), a BAC (Bacterial Artificial Chromosome), a phage, a phagemid, a cosmid, a plasmid or even a linear DNA molecule which may consist of a chromosomal, non-chromosomal, semi-synthetic and synthetic DNA. Such a recombinant
- 10 vector can comprise a transcriptional unit comprising an assembly of :
- (1) a genetic element or elements having a regulatory role in gene expression, for example promoters or enhancers. Enhancers are cis-acting elements of DNA, usually from about 10 to 300 bp in length that act on the promoter to increase the transcription.
  - (2) a structural or coding sequence which is transcribed into mRNA and eventually
  - 15 translated into a polypeptide, said structural or coding sequence being operably linked to the regulatory elements described in (1); and
  - (3) appropriate transcription initiation and termination sequences. Structural units intended for use in yeast or eukaryotic expression systems preferably include a leader sequence enabling extracellular secretion of translated protein by a host cell. Alternatively, when a
  - 20 recombinant protein is expressed without a leader or transport sequence, it may include a N-terminal residue. This residue may or may not be subsequently cleaved from the expressed recombinant protein to provide a final product.

Generally, recombinant expression vectors will include origins of replication, selectable markers permitting transformation of the host cell, and a promoter derived from a

25 highly expressed gene to direct transcription of a downstream structural sequence. The heterologous structural sequence is assembled in appropriate phase with translation initiation and termination sequences, and preferably a leader sequence capable of directing secretion of the translated protein into the periplasmic space or the extracellular medium. In a specific embodiment wherein the vector is adapted for transfecting and expressing desired

30 sequences in mammalian host cells, preferred vectors will comprise an origin of replication in the desired host, a suitable promoter and enhancer, and also any necessary ribosome binding sites, polyadenylation site, splice donor and acceptor sites, transcriptional termination sequences, and 5'-flanking non-transcribed sequences. DNA sequences derived

from the SV40 viral genome, for example SV40 origin, early promoter, enhancer, splice and polyadenylation sites may be used to provide the required non-transcribed genetic elements.

The *in vivo* expression of a MGST-II polypeptide of SEQ ID Nos. 488 and 489 may be useful in order to correct a genetic defect related to the expression of the native gene in a host organism or to the production of a biologically inactive MGST-II protein.

Consequently, the present invention also deals with recombinant expression vectors mainly designed for the *in vivo* production of the MGST-II polypeptide of SEQ ID Nos. 488 and 489 or fragments or variants thereof by the introduction of the appropriate genetic material in the organism of the patient to be treated. This genetic material may be introduced *in vitro* in a cell that has been previously extracted from the organism, the modified cell being subsequently reintroduced in the said organism, directly *in vivo* into the appropriate tissue.

## 2. Regulatory elements

### Promoters:

The suitable promoter regions used in the expression vectors according to the present invention are chosen taking into account the cell host in which the heterologous gene has to be expressed. The particular promoter employed to control the expression of a nucleic acid sequence of interest is not believed to be important, so long as it is capable of directing the expression of the nucleic acid in the targeted cell. Thus, where a human cell is targeted, it is preferable to position the nucleic acid coding region adjacent to and under the control of a promoter that is capable of being expressed in a human cell, such as, for example, a human or a viral promoter.

A suitable promoter may be heterologous with respect to the nucleic acid for which it controls the expression or alternatively can be endogenous to the native polynucleotide containing the coding sequence to be expressed. Additionally, the promoter is generally heterologous with respect to the recombinant vector sequences within which the construct promoter/coding sequence has been inserted.

Promoter regions can be selected from any desired gene using, for example, CAT (chloramphenicol transferase) vectors and more preferably pKK232-8 and pCM7 vectors.

Preferred bacterial promoters are the LacI, LacZ, the T3 or T7 bacteriophage RNA polymerase promoters, the gpt, lambda PR, PL and trp promoters (EP 0036776), the polyhedrin promoter, or the p10 protein promoter from baculovirus (Kit Novagen) (Smith et al., 1983; O'Reilly et al., 1992), the lambda PR promoter or also the trc promoter.

Eukaryotic promoters include CMV immediate early, HSV thymidine kinase, early and late SV40, LTRs from retrovirus, and mouse metallothionein-L. Selection of a convenient vector and promoter is well within the level of ordinary skill in the art.

The choice of a promoter is well within the ability of a person skilled in the field of genetic engineering. For example, one may refer to the book of Sambrook et al. (*Molecular Cloning: A Laboratory Manual*, 2nd edition, Cold Spring Harbor Laboratory, Cold Spring Harbor, New York, 1989).

Other regulatory elements:

Where a cDNA insert is employed, one will typically desire to include a polyadenylation signal to effect proper polyadenylation of the gene transcript. The nature of the polyadenylation signal is not believed to be crucial to the successful practice of the invention, and any such sequence may be employed such as human growth hormone and SV40 polyadenylation signals. Also contemplated as an element of the expression cassette is a terminator. These elements can serve to enhance message levels and to minimize read through from the cassette into other sequences.

The vector containing the appropriate DNA sequence as described above, more preferably MGST-II gene regulatory polynucleotide, a polynucleotide encoding the MGST-II polypeptides of SEQ ID Nos. 488 and 489 or both of them, can be utilized to transform an appropriate host to allow the expression of the desired polypeptide or polynucleotide.

### 3. Selectable markers

Such markers would confer an identifiable change to the cell permitting easy identification of cells containing the expression construct. The selectable marker genes for selection of transformed host cells are preferably dihydrofolate reductase or neomycin resistance for eukaryotic cell culture, TRP1 for *S. cerevisiae* or tetracycline, rifampicin or ampicillin resistance in *E. coli*, or levan saccharase for mycobacteria, this latter marker being a negative selection marker.

### 4. Preferred vectors

Bacterial vectors:

As a representative but non-limiting example, useful expression vectors for bacterial use can comprise a selectable marker and a bacterial origin of replication derived from commercially available plasmids comprising genetic elements of pBR322 (ATCC 37017). Such commercial vectors include, for example, pKK223-3 (Pharmacia, Uppsala, Sweden), and GEM1 (Promega Biotec, Madison, WI, USA). Large numbers of other suitable vectors are known to those of skill in the art, and commercially available, such as the following

bacterial vectors : pQE70, pQE60, pQE-9 (Qiagen), pbs, pD10, phagescript, psiX174, pbluescript SK, pbsks, pNH8A, pNH16A, pNH18A, pNH46A (Stratagene); ptrc99a, pKK223-3, pKK233-3, pDR540, pRIT5 (Pharmacia); pWLNEO, pSV2CAT, pOG44, pXT1, pSG (Stratagene); pSVK3, pBPV, pMSG, pSVL (Pharmacia); pQE-30 (QIAexpress).

## 5 Bacteriophage vectors

The P1 bacteriophage vector may contain large inserts ranging from about 80 to about 100 kb. The construction of P1 bacteriophage vectors such as p158 or p158/neo8 have been described by Sternberg (*Mamm. Genome*, 5:397-404, 1994). Recombinant P1 clones comprising MGST-II nucleotide sequences may be designed for inserting large  
 10 polynucleotides of more than 40 kb (Linton et al., *J. Clin. Invest.*, 92:3029-3037, 1993). To generate P1 DNA for transgenic experiments, a preferred protocol is the protocol described by McCormick et al. (*Genet. Anal. Tech. Appl.*, 11:158-164, 1994). Briefly, *E. coli* (preferably strain NS3529) harboring the P1 plasmid are grown overnight in a suitable broth medium containing 25 µg/ml of kanamycin. The P1 DNA is prepared from the *E. coli* by  
 15 alkaline lysis using the Qiagen Plasmid Maxi kit (Qiagen, Chatsworth, CA, USA), according to the manufacturer's instructions. The P1 DNA is purified from the bacterial lysate on two Qiagen-tip 500 columns, using the washing and elution buffers contained in the kit. A phenol/chloroform extraction is then performed before precipitating the DNA with 70% ethanol. After solubilizing the DNA in TE (10 mM Tris-HCl, pH 7.4, 1 mM  
 20 EDTA), the concentration of the DNA is assessed by spectrophotometry.

When the goal is to express a P1 clone comprising MGST-II nucleotide sequences in a transgenic animal, typically in transgenic mice, it is desirable to remove vector sequences from the P1 DNA fragment, for example by cleaving the P1 DNA at rare-cutting sites within the P1 polylinker (*SfiI*, *NotI* or *SalI*). The P1 insert is then purified from vector  
 25 sequences on a pulsed-field agarose gel, using methods similar using methods similar to those originally reported for the isolation of DNA from YACs (Schedl et al., 1993a; Peterson et al., 1993). At this stage, the resulting purified insert DNA can be concentrated, if necessary, on a Millipore Ultrafree-MC Filter Unit (Millipore, Bedford, MA, USA – 30,000 molecular weight limit) and then dialyzed against microinjection buffer (10 mM  
 30 Tris-HCl, pH 7.4; 250 µM EDTA) containing 100 mM NaCl, 30 µM spermine, 70 µM spermidine on a microdialysis membrane (type VS, 0.025 µM from Millipore). The intactness of the purified P1 DNA insert is assessed by electrophoresis on 1% agarose (Sea Kem GTG; FMC Bio-products) pulse-field gel and staining with ethidium bromide.

Baculovirus vectors:

A suitable vector for the expression of the MGST-II polypeptides of SEQ ID Nos. 488 and 489 is a baculovirus vector that can be propagated in insect cells and in insect cell lines. A specific suitable host vector system is the pVL1392/1393 baculovirus transfer vector (Pharmingen) that is used to transfect the SF9 cell line (ATCC N<sup>o</sup>CRL 1711) which  
5 is derived from *Spodoptera frugiperda*.

Other suitable vectors for the expression of the MGST-II polypeptides of SEQ ID Nos. 488 and 489 in a baculovirus expression system include those described by Chai et al. (*Biotech. Appl. Biochem.*, 18:259-273, 1993), Vlasak et al. (*Eur. J. Biochem.*, 135: 123-126, 1983) and Lenhard et al. (*Gene*, 169: 187-190, 1996).

#### 10 Viral vectors

Retrovirus vectors and adeno-associated virus vectors are generally understood to be the recombinant gene delivery systems of choice for the transfer of exogenous polynucleotides *in vivo*, particularly to mammals, including humans. These vectors provide efficient delivery of genes into cells, and the transferred nucleic acids are stably integrated  
15 into the chromosomal DNA of the host.

Particularly preferred retroviruses for the preparation or construction of retroviral *in vitro* or *in vitro* gene delivery vehicles of the present invention include retroviruses selected from the group consisting of Mink-Cell Focus Inducing Virus, Murine Sarcoma Virus, Reticuloendotheliosis virus and Rous Sarcoma virus. Particularly preferred Murine  
20 Leukemia Viruses include the 4070A and the 1504A viruses, Abelson (ATCC No VR-999), Friend (ATCC No VR-245), Gross (ATCC No. VR-590), Rauscher (ATCC No VR-998) and Moloney Murine Leukemia Virus (ATCC No VR-190; PCT Application No WO 94/24298). Particularly preferred Rous Sarcoma Viruses include Bryan high titer (ATCC Nos. VR-334, VR-657, VR-726, VR-659 and VR-728). Other preferred retroviral vectors  
25 are those described in Roth et al. (*Nature Medicine*, 2:985-991, 1996), PCT Application No. WO 93/25234 and PCT Application No. WO 94/ 06920.

Yet another viral vector system that is contemplated by the invention consists in the adeno-associated virus (AAV). The adeno-associated virus is a naturally occurring defective virus that requires another virus, such as an adenovirus or a herpes virus, as a helper virus  
30 for efficient replication and a productive life cycle (Muzyczka et al., *Current Topics in Microbiol. Immunol.*, 158:97-129, 1992). It is also one of the few viruses that may integrate its DNA into non-dividing cells, and exhibits a high frequency of stable integration (McLaughlin et al., *Am. J. Hum. Genet.*, 59: 561-569, 1989). One advantageous feature of

AAV derives from its reduced efficacy for transducing primary cells relative to transformed cells.

BAC vectors:

The bacterial artificial chromosome (BAC) cloning system (Shizuya et al., 1992) has been developed to stably maintain large fragments of genomic DNA (100-300 kb) in *E. coli*. A preferred BAC vector consists of pBeloBAC11 vector that has been described by Kim et al. (*Genomics*, 34:213-218,1996). BAC libraries are prepared with this vector using size-selected genomic DNA that has been partially digested using enzymes that permit ligation into either the *Bam* HI or *Hind*III sites in the vector. Flanking these cloning sites are T7 and SP6 RNA polymerase transcription initiation sites that can be used to generate end probes by either RNA transcription or PCR methods. After the construction of a BAC library in *E. coli*, BAC DNA is purified from the host cell as a supercoiled circle. Converting these circular molecules into a linear form precedes both size determination and introduction of the BACs into recipient cells. The cloning site is flanked by two *Not* I sites, permitting cloned segments to be excised from the vector by *Not* I digestion. Alternatively, the DNA insert contained in the pBeloBAC11 vector may be linearized by treatment of the BAC vector with the commercially available enzyme lambda terminase that leads to the cleavage at the unique *cos*N site, but this cleavage method results in a full length BAC clone containing both the insert DNA and the BAC sequences.

#### 5. Delivery of the recombinant vectors

In order to effect expression of the polynucleotides and polynucleotide constructs of the invention, these constructs must be delivered into a cell. This delivery may be accomplished *in vitro*, as in laboratory procedures for transforming cell lines, or *in vivo* or *ex vivo*, as in the treatment of certain diseases states. One mechanism is viral infection where the expression construct is encapsidated in an infectious viral particle. Several non-viral methods for the transfer of polynucleotides into cultured mammalian cells are also contemplated by the present invention, and include, without being limited to, calcium phosphate precipitation (Chen et al., *Proc. Natl. Acad. Sci. USA*, 94:10756-10761, 1987), DEAE-dextran (Gopal, *Mol. Cell. Biol.*, 5:1188-1190, 1985), electroporation (Tur-Kaspa et al., *Mol. Cell. Biol.*, 6:716-718, 1986), direct microinjection (Harland et al., 1985), DNA-loaded liposomes (Nicolau et al., 1982; Fraley et al., 1979), and receptor-mediate transfection (Wu and Wu, 1987; 1988). Some of these techniques may be successfully adapted for *in vivo* or *ex vivo* use.

Once the expression polynucleotide has been delivered into the cell, it may be stably integrated into the genome of the recipient cell. This integration may be in the cognate location and orientation via homologous recombination (gene replacement) or it may be integrated in a random, non-specific location (gene augmentation). In yet further  
5   embodiments, the nucleic acid may be stably maintained in the cell as a separate, episomal segment of DNA. Such nucleic acid segments or "episomes" encode sequences sufficient to permit maintenance and replication independent of or in synchronization with the host cell cycle.

One specific embodiment for a method for delivering a protein or peptide to the  
10   interior of a cell of a vertebrate *in vivo* comprises the step of introducing a preparation comprising a physiologically acceptable carrier and a naked polynucleotide operatively coding for the polypeptide of interest into the interstitial space of a tissue comprising the cell, whereby the naked polynucleotide is taken up into the interior of the cell and has a physiological effect. This is particularly applicable for transfer *in vitro* but it may be applied  
15   to *in vivo* as well.

Compositions for use *in vitro* and *in vivo* comprising a "naked" polynucleotide are described in PCT application No. WO 90/11092 (Vical Inc.) and also in PCT application No. WO 95/11307.

In still another embodiment of the invention, the transfer of a naked polynucleotide  
20   of the invention, including a polynucleotide construct of the invention, into cells may be proceeded with a particle bombardment (biolistic), said particles being DNA-coated microprojectiles accelerated to a high velocity allowing them to pierce cell membranes and enter cells without killing them, such as described by Klein et al. (*Nature* 327:70-73, 1987).

In a further embodiment, the polynucleotide of the invention may be entrapped in a  
25   liposome (Ghosh and Bacchawat, *Targeting of liposomes to hepatocytes*, In: *Liver Diseases, Targeted diagnosis and therapy using specific receptors and ligands*, Marcel Dekeker, New York, 87-104, 1991; Wong et al., *Gene* 10:87-94, 1980; Nicolau et al., *Biochim. Biophys. Acta*. 721:185-190, 1982).

In a specific embodiment, the invention provides a composition for the *in vivo*  
30   production of the MGST-II protein or polypeptide described herein. It comprises a naked polynucleotide operatively coding for this polypeptide, in solution in a physiologically acceptable carrier, and suitable for introduction into a tissue to cause cells of the tissue to express the said protein or polypeptide.



The amount of vector to be injected to the desired host organism varies according to the site of injection. As an indicative dose, it will be injected between 0.1 and 100  $\mu$ g of the vector in an animal body, preferably a mammal body, for example a mouse body.

In another embodiment of the invention, the vector may be introduced *in vitro* in a  
5 host cell, preferably in a host cell previously harvested from the animal to be treated and more preferably a somatic cell such as a muscle cell. In a subsequent step, the cell that has been transformed with the vector coding for the desired MGST-II polypeptide or the desired fragment thereof is reintroduced into the animal body in order to deliver the recombinant protein within the body either locally or systemically.

#### 10 Host Cells

Another embodiment of the invention consists of a host cell that has been transformed or transfected with one of the polynucleotides described therein, and more precisely a polynucleotide either comprising a MGST-II regulatory polynucleotide or the coding sequence of the MGST-II polypeptide having the amino acid sequence of SEQ ID  
15 Nos. 488 and 489. The embodiment includes host cells that are transformed (prokaryotic cells) or that are transfected (eukaryotic cells) with a recombinant vector such as one of those described above. Generally, a recombinant host cell of the invention comprises any one of the polynucleotides or the recombinant vectors described therein.

A preferred recombinant host cell according to the invention comprises a  
20 polynucleotide selected from the following group of polynucleotides :  
a) a purified or isolated nucleic acid encoding a MGST-II polypeptide, or a polypeptide fragment or variant thereof.  
b) a purified or isolated nucleic comprising at least 8, preferably at least 15, more preferably at least 25, consecutive nucleotides of the nucleotide sequence SEQ ID No. 485, a  
25 nucleotide sequence complementary thereto, or a variant thereof.  
c) a purified or isolated nucleic acid comprising at least 8 consecutive nucleotides, preferably at least 15, more preferably at least 25 of the nucleotide sequence SEQ ID Nos. 486 and 487, a nucleotide sequence complementary thereto or a variant thereof.  
d) a purified or isolated nucleic acid comprising an exon of the MGST-II gene, a sequence  
30 complementary thereto or a fragment or a variant thereof.  
e) a purified or isolated nucleic acid comprising a combination of at least two exons of the MGST-II gene, or the sequences complementary thereto wherein the polynucleotides are arranged within the nucleic acid, from the 5' end to the 3' end of said nucleic acid, in the same order than in SEQ ID No. 485.

- f) a purified or isolated nucleic acid comprising the nucleotide sequence SEQ ID No. 485 or the sequences complementary thereto or a biologically active fragment thereof.
- g) a purified or isolated nucleic acid comprising the nucleotide sequence SEQ ID No. 486, or the sequence complementary thereto or a biologically active fragment thereof.
- 5 h) a polynucleotide consisting of:
- (1) a nucleic acid comprising a regulatory polynucleotide of SEQ ID No. 485 or the sequences complementary thereto or a biologically active fragment thereof
  - (2) a polynucleotide encoding a desired polypeptide or nucleic acid.
  - (3) Optionally, a nucleic acid comprising a regulatory polynucleotide of SEQ ID No. 485, or
- 10 the sequence complementary thereto or a biologically active fragment thereof.
- i) a DNA construct as described previously in the present specification.

Another preferred recombinant cell host according to the present invention is characterized in that its genome or genetic background (including chromosome, plasmids) is modified by the nucleic acid coding for the MGST-II polypeptide of SEQ ID Nos. 488

15 and 489 or fragments or variants thereof.

Preferred host cells used as recipients for the expression vectors of the invention are the following:

- a) Prokaryotic host cells: *Escherichia coli* strains (I.E. DH5- $\alpha$  strain), *Bacillus subtilis*, *Salmonella typhimurium*, and strains from species like *Pseudomonas*, *Streptomyces* and
- 20 *Staphylococcus*.
- b) Eukaryotic host cells: HeLa cells (ATCC N°CCL2; N°CCL2.1; N°CCL2.2), Cv 1 cells (ATCC N°CCL70), COS cells (ATCC N°CRL1650; N°CRL1651), Sf-9 cells (ATCC N°CRL1711), C127 cells (ATCC N° CRL-1804), 3T3 (ATCC N° CRL-6361), CHO (ATCC N° CCL-61), human kidney 293.(ATCC N° 45504; N° CRL-1573) and BHK (ECACC N°
- 25 84100501; N° 84111301)
- c) Other mammalian host cells:

The MGST-II gene expression in mammalian, and typically human, cells may be rendered defective, or alternatively it may be proceeded with the insertion of a MGST-II genomic or cDNA sequence with the replacement of the MGST-II gene counterpart in the

30 genome of an animal cell by a MGST-II polynucleotide according to the invention. These genetic alterations may be generated by homologous recombination events using specific DNA constructs that have been previously described.

One kind of cell hosts that may be used are mammal zygotes, such as murine zygotes. For example, murine zygotes may undergo microinjection with a purified DNA

molecule of interest, for example a purified DNA molecule that has previously been adjusted to a concentration range from 1 ng/ml (for BAC inserts) 3 ng/ $\mu$ l (for P1 bacteriophage inserts) in 10 mM Tris-HCl, pH 7.4, 250  $\mu$ M EDTA containing 100 mM NaCl, 30  $\mu$ M spermine, and 70  $\mu$ M spermidine. When the DNA to be microinjected has a large size, polyamines and high salt concentrations can be used in order to avoid mechanical breakage of this DNA, as described by Schedl et al (*Nucleic Acids Res.* 21:4783-4787, 1993).

Anyone of the polynucleotides of the invention, including the DNA constructs described herein, may be introduced in an embryonic stem (ES) cell line, preferably a mouse ES cell line. ES cell lines are derived from pluripotent, uncommitted cells of the inner cell mass of pre-implantation blastocysts. Preferred ES cell lines are the following: ES-E14TG2a (ATCC n° CRL-1821), ES-D3 (ATCC n° CRL1934 and n° CRL-11632), YS001 (ATCC n° CRL-11776), 36.5 (ATCC n° CRL-11116). To maintain ES cells in an uncommitted state, they are cultured in the presence of growth inhibited feeder cells which, provide the appropriate signals to preserve this embryonic phenotype and serve as a matrix for ES cell adherence. Preferred feeder cells consist of primary embryonic fibroblasts that are established from tissue of day 13- day 14 embryos of virtually any mouse strain, that are maintained in culture, such as described by Abbondanzo et al. (*Methods in Enzymology*, Academic Press, New York, 803-823, 1993) and are inhibited in growth by irradiation, such as described by Robertson ("Embryo-Derived Stem Cell Lines," *E.J. Robertson Ed. Teratocarcinomas and Embryonic Stem Cells: A Practical Approach*. IRL Press, Oxford, 71, 1987), or by the presence of an inhibitory concentration of LIF, such as described by Pease and Williams (*Exp. Cell. Res.* 190:09-211, 1990).

The constructs in the host cells can be used in a conventional manner to produce the gene product encoded by the recombinant sequence.

Following transformation of a suitable host and growth of the host to an appropriate cell density, the selected promoter is induced by appropriate means, such as temperature shift or chemical induction, and cells are cultivated for an additional period.

Cells are typically harvested by centrifugation, disrupted by physical or chemical means, and the resulting crude extract retained for further purification.

Microbial cells employed in the expression of proteins can be disrupted by any convenient method, including freeze-thaw cycling, sonication, mechanical disruption, or use of cell lysing agents. Such methods are well known by one skilled in the art.

### Transgenic animals

The terms "transgenic animals" or "host animals" are used herein designate animals that have their genome genetically and artificially manipulated so as to include one of the nucleic acids according to the invention. Preferred animals are non-human mammals and  
5 include those belonging to a genus selected from *Mus* (e.g. mice), *Rattus* (e.g. rats) and *Oryctogalus* (e.g. rabbits) which have their genome artificially and genetically altered by the insertion of a nucleic acid according to the invention.

The transgenic animals of the invention all include within a plurality of their cells a cloned recombinant or synthetic DNA sequence, more specifically one of the purified or  
10 isolated nucleic acids comprising a MGST-II coding sequence, a MGST-II regulatory polynucleotide or a DNA sequence encoding an antisense polynucleotide such as described in the present specification.

Preferred transgenic animals according to the invention contains in their somatic cells and/or in their germ line cells a polynucleotide selected from the following group of  
15 polynucleotides :

- a) a purified or isolated nucleic acid encoding a MGST-II polypeptide, or a polypeptide fragment or variant thereof.
- b) a purified or isolated nucleic comprising at least 8, preferably at least 15, more preferably at least 25, consecutive nucleotides of the nucleotide sequence SEQ ID No. 485, a  
20 nucleotide sequence complementary thereto.
- c) a purified or isolated nucleic acid comprising at least 8 consecutive nucleotides, preferably at least 15, more preferably at least 25 of the nucleotide sequence SEQ ID Nos. 486 and 487, a nucleotide sequence complementary thereto.
- d) a purified or isolated nucleic acid comprising an exon of the MGST-II gene, a sequence  
25 complementary thereto or a fragment or a variant thereof.
- e) a purified or isolated nucleic acid comprising a combination of at least two exons of the MGST-II gene, or the sequences complementary thereto wherein the polynucleotides are arranged within the nucleic acid, from the 5' end to the 3' end of said nucleic acid, in the same order than in SEQ ID No. 485.
- 30 f) a purified or isolated nucleic acid comprising the nucleotide sequence SEQ ID No. 485 or the sequences complementary thereto or a biologically active fragment thereof.
- g) a purified or isolated nucleic acid comprising the nucleotide sequence SEQ ID No. 486, or the sequence complementary thereto or a biologically active fragment thereof.
- h) a polynucleotide consisting of :

- (1) a nucleic acid comprising a regulatory polynucleotide of SEQ ID No. 485 or the sequences complementary thereto or a biologically active fragment thereof
- (2) a polynucleotide encoding a desired polypeptide or nucleic acid.
- (3) Optionally, a nucleic acid comprising a regulatory polynucleotide of SEQ ID No. 486, or
- 5 the sequence complementary thereto or a biologically active fragment thereof.
- i) a DNA construct as described previously in the present specification.

The transgenic animals of the invention thus contain specific sequences of exogenous genetic material such as the nucleotide sequences described above in detail.

In a first preferred embodiment, these transgenic animals may be good experimental

10 models in order to study the diverse pathologies related to cell differentiation, in particular concerning the transgenic animals within the genome of which has been inserted one or several copies of a polynucleotide encoding a native MGST-II protein, or alternatively a mutant MGST-II protein.

In a second preferred embodiment, these transgenic animals may express a desired

15 polypeptide of interest under the control of the regulatory polynucleotides of the MGST-II gene, leading to good yields in the synthesis of this protein of interest, and eventually a tissue specific expression of this protein of interest.

The design of the transgenic animals of the invention may be made according to the conventional techniques well known from the one skilled in the art. For more details

20 regarding the production of transgenic animals, and specifically transgenic mice, it may be referred to US Patents Nos. 4,873,191, issued October 10, 1989, 5,464,764 issued November 7, 1995 and 5,789,215, issued August 4, 1998, these documents being herein incorporated by reference to disclose methods producing transgenic mice.

Transgenic animals of the present invention are produced by the application of

25 procedures which result in an animal with a genome that has incorporated exogenous genetic material. The procedure involves obtaining the genetic material, or a portion thereof, which encodes either a MGST-II coding sequence, a MGST-II regulatory polynucleotide or a DNA sequence encoding a MGST-II antisense polynucleotide such as described in the present specification.

30 A recombinant polynucleotide of the invention is inserted into an embryonic or ES stem cell line. The insertion is preferably made using electroporation, such as described by Thomas et al. (*Cell* 51:503-512, 1987). The cells subjected to electroporation are screened (e.g. by selection via selectable markers, by PCR or by Southern blot analysis) to find positive cells which have integrated the exogenous recombinant polynucleotide into their

genome, preferably via an homologous recombination event. An illustrative positive-negative selection procedure that may be used according to the invention is described by Mansour et al. (*Nature* 336:348-352, 1988).

Then, the positive cells are isolated, cloned and injected into 3.5 days old blastocysts from mice, such as described by Bradley ("Production and Analysis of Chimaeric Mice," *E.J. Robertson (Ed.), Teratocarcinomas and embryonic stem cells: A practical approach* IRL Press, Oxford, 113, 1987). The blastocysts are then inserted into a female host animal and allowed to grow to term.

Alternatively, the positive ES cells are brought into contact with embryos at the 2.5 days old 8-16 cell stage (morulae) such as described by Wood et al. (*Proc. Natl. Acad. Sci. U.S.A.* 90:4582-4585, 1993) or by Nagy et al. (*Proc. Natl. Acad. Sci. USA.* 90: 8424-8428, 1993), the ES cells being internalized to colonize extensively the blastocyst including the cells which will give rise to the germ line.

The offspring of the female host are tested to determine which animals are transgenic e.g. include the inserted exogenous DNA sequence and which are wild-type.

Thus, the present invention also concerns a transgenic animal containing a nucleic acid, a recombinant expression vector or a recombinant host cell according to the invention.

A further object of the invention consists of recombinant host cells obtained from a transgenic animal described herein.

Recombinant cell lines may be established *in vitro* from cells obtained from any tissue of a transgenic animal according to the invention, for example by transfection of primary cell cultures with vectors expressing *onc*-genes such as SV40 large T antigen, as described by Chou (*Mol. Endocrinol.* 3:1511-1514, 1989) and Shay et al. (*Biochem. Biophys. Acta.* 1072:1-7, 1991).

## **I.E. MGST-II Polypeptides**

The term "MGST-II polypeptides" is used herein to embrace all of the proteins and polypeptides of the present invention. Also forming part of the invention are polypeptides encoded by the polynucleotides of the invention, as well as fusion polypeptides comprising such polypeptides. The invention embodies MGST-II proteins from humans, including isolated or purified MGST-II proteins consisting, consisting essentially, or comprising the sequence of SEQ ID Nos. 488 and 489. It should be noted the MGST-II proteins of the invention are based on the naturally-occurring variants of the amino acid sequence of human MGST-II.

In a first embodiment, the present invention provides a variant MGST-II protein; wherein the Tyr residue of amino acid position 93 has been replaced with a His residue. Variant proteins and the fragments thereof which contain amino acid position 93 are collectively referred to herein as "93-His variants." More particularly, the present invention  
5 embodies isolated, purified, and recombinant polypeptides comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, 30, 40, 50, or 100 amino acids of SEQ ID No. 488, wherein said contiguous span comprises a His residue at amino acid position 93. In this amino acid substitution the original residue (Tyr) is replaced by a non-equivalent amino acid (His) presenting different  
10 chemical properties.

The present invention further provides another naturally-occurring variant of the MGST-II protein that consists or consists essentially of amino acids 1-109 of SEQ ID No. 488. This variant MGST-II polypeptide corresponds to one allele of biallelic marker 10-290-37.

15 Another naturally-occurring variant of the MGST-II protein of the present invention is encoded by a cDNA obtained by alternative splicing. MGST-II cDNAs and cDNA variants are further described above. This variant polypeptide of a sequence from SEQ ID No. 489 is identical to the MGST-II protein of SEQ ID No. 488 from amino acid position 1 to amino acid position 19 but comprises 11 additional amino acids. The present invention  
20 embodies isolated, purified, and recombinant polypeptides comprising, consisting of or consisting essentially of an amino acid sequence from SEQ ID No. 489. Moreover, the present invention embodies isolated, purified, and recombinant polypeptides comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, or 30 amino acids of SEQ ID No. 489, wherein said  
25 contiguous span comprises a least one of amino acid positions 20 to 30 of SEQ ID No. 489.

All the variant MGST-II polypeptides described above most probably show alterations in the activity, specificity and function of the MGST-II enzyme. In preferred embodiments the polypeptides of the present invention comprise the site of a mutation or functional mutation, including a deletion, substitution or truncation in the amino acid  
30 sequence in the MGST-II protein.

MGST-II proteins are preferably isolated from human or mammalian tissue samples or expressed from human or mammalian genes. The MGST-II polypeptides of the invention can be made using routine expression methods known in the art. The polynucleotide encoding the desired polypeptide, is ligated into an expression vector suitable for any

convenient host. Both eukaryotic and prokaryotic host systems are used in forming recombinant polypeptides. The polypeptide is then isolated from lysed cells or from the culture medium and purified to the extent needed for its intended use. Purification is by any technique known in the art, for example, differential extraction, salt fractionation, chromatography, centrifugation, and the like. See, for example, Methods in Enzymology for a variety of methods for purifying proteins.

In addition, shorter protein fragments are produced by chemical synthesis. Alternatively the proteins of the invention are extracted from cells or tissues of humans or non-human animals. Methods for purifying proteins are known in the art, and include the use of detergents or chaotropic agents to disrupt particles followed by differential extraction and separation of the polypeptides by ion exchange chromatography, affinity chromatography, sedimentation according to density, and gel electrophoresis.

Any MGST-II cDNA of the invention is used to express MGST-II proteins and polypeptides. The nucleic acid encoding the MGST-II protein or polypeptide to be expressed is operably linked to a promoter in an expression vector using conventional cloning technology. The MGST-II insert in the expression vector may comprise the full coding sequence for the MGST-II protein or a portion thereof. For example, the MGST-II derived insert may encode a polypeptide comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, 30, 40, 50, or 100 amino acids of SEQ ID No. 488, wherein said contiguous span comprises a His residue at amino acid position 93. The MGST-II derived insert may further encode a polypeptide comprising, consisting of or consisting essentially of an amino acid sequence from amino acid positions 1-108 of SEQ ID No. 488. The MGST-II derived insert may further encode a polypeptide comprising, consisting of or consisting essentially of an amino acid sequence from SEQ ID No. 489. The MGST-II derived insert may also encode a polypeptide comprising a contiguous span of at least 6 amino acids, preferably at least 8 to 10 amino acids, more preferably at least 12, 15, 20, 25, or 30 amino acids of SEQ ID No. 489, wherein said contiguous span comprises a least one of amino acid positions 20 to 30 of SEQ ID No. 489.

The expression vector is any of the mammalian, yeast, insect or bacterial expression systems known in the art. Commercially available vectors and expression systems are available from a variety of suppliers including Genetics Institute (Cambridge, MA), Stratagene (La Jolla, California), Promega (Madison, Wisconsin), and Invitrogen (San Diego, California). If desired, to enhance expression and facilitate proper protein folding, the codon context and



codon pairing of the sequence is optimized for the particular expression organism in which the expression vector is introduced, as explained by Hatfield, et al., U.S. Patent No. 5,082,767.

In one embodiment, the entire coding sequence of the MGST-II cDNA through the poly A signal of the cDNA is operably linked to a promoter in the expression vector.

- 5 Alternatively, if the nucleic acid encoding a portion of the MGST-II protein lacks a methionine to serve as the initiation site, an initiating methionine can be introduced next to the first codon of the nucleic acid using conventional techniques. Similarly, if the insert from the MGST-II cDNA lacks a poly A signal, this sequence can be added to the construct by, for example, splicing out the Poly A signal from pSG5 (Stratagene) using BglI and SalI restriction
- 10 endonuclease enzymes and incorporating it into the mammalian expression vector pXT1 (Stratagene). pXT1 contains the LTRs and a portion of the gag gene from Moloney Murine Leukemia Virus. The position of the LTRs in the construct allow efficient stable transfection. The vector includes the Herpes Simplex Thymidine Kinase promoter and the selectable neomycin gene. The nucleic acid encoding the MGST-II protein or a portion thereof is
- 15 obtained by PCR from a bacterial vector containing a MGST-II cDNA of the present invention using oligonucleotide primers complementary to the MGST-II cDNA or portion thereof and containing restriction endonuclease sequences for Pst I incorporated into the 5' primer and BglII at the 5' end of the corresponding cDNA 3' primer, taking care to ensure that the sequence encoding the MGST-II protein or a portion thereof is positioned properly with respect
- 20 to the poly A signal. The purified fragment obtained from the resulting PCR reaction is digested with PstI, blunt ended with an exonuclease, digested with Bgl II, purified and ligated to pXT1, now containing a poly A signal and digested with BglII.

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc., Grand Island, New York) under conditions outlined in the product

25 specification. Positive transfectants are selected after growing the transfected cells in 600ug/ml G418 (Sigma, St. Louis, Missouri).

Alternatively, the nucleic acids encoding the MGST-II protein or a portion thereof is cloned into pED6dpc2 (Genetics Institute, Cambridge, MA). The resulting pED6dpc2 constructs is transfected into a suitable host cell, such as COS 1 cells. Methotrexate resistant

30 cells are selected and expanded.

The above procedures may also be used to express a mutant MGST-II protein responsible for a detectable phenotype or a portion thereof.

The expressed proteins are purified using conventional purification techniques such as ammonium sulfate precipitation or chromatographic separation based on size or charge. The

protein encoded by the nucleic acid insert may also be purified using standard immunochromatography techniques. In such procedures, a solution containing the expressed MGST-II protein or portion thereof, such as a cell extract, is applied to a column having antibodies against the MGST-II protein or portion thereof is attached to the chromatography matrix. The expressed protein is allowed to bind the immunochromatography column. Thereafter, the column is washed to remove non-specifically bound proteins. The specifically bound expressed protein is then released from the column and recovered using standard techniques.

To confirm expression of the MGST-II protein or a portion thereof, the proteins expressed from host cells containing an expression vector containing an insert encoding the MGST-II protein or a portion thereof can be compared to the proteins expressed in host cells containing the expression vector without an insert. The presence of a band in samples from cells containing the expression vector with an insert which is absent in samples from cells containing the expression vector without an insert indicates that the MGST-II protein or a portion thereof is being expressed. Generally, the band will have the mobility expected for the MGST-II protein or portion thereof. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

Antibodies capable of specifically recognizing the expressed MGST-II protein or a portion thereof, are described below.

If antibody production is not possible, the nucleic acids encoding the MGST-II protein or a portion thereof is incorporated into expression vectors designed for use in purification schemes employing chimeric polypeptides. In such strategies the nucleic acid encoding the MGST-II protein or a portion thereof is inserted in frame with the gene encoding the other half of the chimera. The other half of the chimera is  $\beta$ -globin or a nickel binding polypeptide encoding sequence. A chromatography matrix having antibody to  $\beta$ -globin or nickel attached thereto is then used to purify the chimeric protein. Protease cleavage sites is engineered between the  $\beta$ -globin gene or the nickel binding polypeptide and the MGST-II protein or portion thereof. Thus, the two polypeptides of the chimera are separated from one another by protease digestion.

One useful expression vector for generating  $\beta$ -globin chimerics is pSG5 (Stratagene), which encodes rabbit  $\beta$ -globin. Intron II of the rabbit  $\beta$ -globin gene facilitates splicing of the expressed transcript, and the polyadenylation signal incorporated into the construct increases the level of expression. These techniques are well known to those skilled in the art of

molecular biology. Standard methods are published in methods texts such as Davis et al., (Basic Methods in Molecular Biology, L.G. Davis, M.D. Digner, and J.F. Battey, ed., Elsevier Press, NY, 1986) and many of the methods are available from Stratagene, Life Technologies, Inc., or Promega. Polypeptide may additionally be produced from the construct using in vitro translation systems such as the In vitro Express™ Translation Kit (Stratagene).

#### **I.F. Production of Antibodies Against MGST-II Polypeptides**

Any MGST-II polypeptide or whole protein may be used to generate antibodies capable of specifically binding to expressed MGST-II protein or fragments thereof or variants thereof. Preferably the antibody compositions of the invention are capable of specifically binding to the 93-His variant of the MGST-II protein. Alternatively the antibody compositions of the present invention are capable of specifically binding the variant MGST-II polypeptide of SEQ ID No. 489. A preferred embodiment of the invention encompasses isolated or purified antibody compositions capable of selectively binding, or which are capable of binding to an epitope-containing fragment of a polypeptide of the invention, wherein said epitope comprises at least one amino acid position selected from the group consisting of His residue at amino acid position 93 of SEQ ID No. 488 and of amino acid positions 20-30 of SEQ ID No. 489. For an antibody composition to specifically bind to these MGST-II variants it must demonstrate at least a 5%, 10%, 15%, 20%, 25%, 50%, or 100% greater binding affinity for full length MGST-II variants in an ELISA, RIA, or other antibody-based binding assay than to full length MGST-II protein described in SEQ ID No. 488. Affinity of the antibody composition for the epitope can further be determined by preparing competitive binding curves, as described, for example, by Fisher, D. (Chap. 42 in: Manual of Clinical Immunology, 2d Ed. (Rose and Friedman, Eds.) *Amer. Soc. For Microbiol.*, Washington, D.C., 1980).

The present invention also contemplates the use of variant MGST-II polypeptides in the manufacture of antibodies. In a preferred embodiment such polypeptides are useful in the manufacture of antibodies to detect the presence and absence of the 93-His variant and of the MGST-II variant of SEQ ID No. 489.

Non-human animals or mammals, whether wild-type or transgenic, which express a different species of MGST-II than the one to which antibody binding is desired, and animals which do not express MGST-II (i.e. an MGST-II knock out animal as described in herein) are particularly useful for preparing antibodies. MGST-II knock out animals will recognize all or most of the exposed regions of MGST-II as foreign antigens, and therefore produce antibodies with a wider array of MGST-II epitopes. Moreover, smaller polypeptides with

only 10 to 30 amino acids may be useful in obtaining specific binding to the 93-His variant and to the MGST-II variant of SEQ ID No. 489. In addition, the humoral immune system of animals which produce a species of MGST-II that resembles the antigenic sequence will preferentially recognize the differences between the animal's native MGST-II species and the antigen sequence, and produce antibodies to these unique sites in the antigen sequence. Such a technique will be particularly useful in obtaining antibodies that specifically bind to the 93-His variant and to the MGST-II variant of SEQ ID No. 489. The preparation of antibody compositions is further described in Example 6.

Antibody preparations prepared according to the present invention are useful in quantitative immunoassays which determine concentrations of antigen-bearing substances in biological samples; they are also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample. The antibodies may also be used in therapeutic compositions for killing cells expressing the protein or reducing the levels of the protein in the body. The antibodies of the invention may be labeled, either by a radioactive, a fluorescent or an enzymatic label. Consequently, the invention is also directed to a method for detecting specifically the presence of a variant MGST-II polypeptide according to the invention in a biological sample, said method comprising the following steps : a) bringing into contact the biological sample with a polyclonal or monoclonal antibody that specifically binds a variant MGST-II polypeptide or to a peptide fragment or variant thereof; and b) detecting the antigen-antibody complex formed. The invention also concerns a diagnostic kit for detecting *in vitro* the presence of a variant MGST-II polypeptide according to the present invention in a biological sample, wherein said kit comprises: a) a polyclonal or monoclonal antibody that specifically binds a variant MGST-II polypeptide or to a peptide fragment or variant thereof, optionally labeled; b) a reagent allowing the detection of the antigen-antibody complexes formed, said reagent carrying optionally a label, or being able to be recognized itself by a labeled reagent, more particularly in the case when the above-mentioned monoclonal or polyclonal antibody is not labeled by itself.

## **II. Methods for *De Novo* Identification of Biallelic Markers**

Large fragments of human DNA, carrying genes of interest involved in the biotransformation of xenobiotics such as therapeutic drugs; were cloned, sequenced and screened for biallelic markers. Biallelic markers within the candidate genes themselves as well as markers located on the same genomic fragment were identified. It will be clear to one of skill in the art that large fragments of human genomic DNA may be obtained from any appropriate source and may be cloned into a number of suitable vectors.

In a preferred embodiment of the invention, BAC (Bacterial Artificial Chromosomes) vectors were used to construct DNA libraries covering the entire human genome. Specific amplification primers were designed for each candidate gene and the BAC library was screened by PCR until there was at least one positive BAC clone per candidate gene. Genomic sequence, screened for biallelic markers, was generated by sequencing ends of BAC subclones. Details of a preferred embodiment are provided in Example 1. As a preferred alternative to sequencing the ends of an adequate number of BAC subclones, high throughput deletion-based sequencing vectors, which allow the generation of a high quality sequence information covering fragments of about 6kb, may be used. Having sequence fragments longer than 2.5 or 3kb enhances the chances of identifying biallelic markers therein. Methods of constructing and sequencing a nested set of deletions are disclosed in the related U.S. Patent Application entitled "High Throughput DNA Sequencing Vector" (Serial No. 09/058,746).

In another embodiment of the invention, genomic sequences of candidate genes were available in public databases allowing direct screening for biallelic markers.

Any of a variety of methods can be used to screen a genomic fragment for single nucleotide polymorphisms such as differential hybridization with oligonucleotide probes, detection of changes in the mobility measured by gel electrophoresis or direct sequencing of the amplified nucleic acid. A preferred method for identifying biallelic markers involves comparative sequencing of genomic DNA fragments from an appropriate number of unrelated individuals.

In a first embodiment, DNA samples from unrelated individuals are pooled together, following which the genomic DNA of interest is amplified and sequenced. The nucleotide sequences thus obtained are then analyzed to identify significant polymorphisms. One of the major advantages of this method resides in the fact that the pooling of the DNA samples substantially reduces the number of DNA amplification reactions and sequencing reactions, which must be carried out. Moreover, this method is sufficiently sensitive so that a biallelic marker obtained thereby usually demonstrates a sufficient frequency of its less common allele to be useful in conducting association studies. Usually, the frequency of the least common allele of a biallelic marker identified by this method is at least 10%.

In a second embodiment, the DNA samples are not pooled and are therefore amplified and sequenced individually. This method is usually preferred when biallelic markers need to be identified in order to perform association studies within candidate genes. Preferably, highly relevant gene regions such as promoter regions or exon regions may be

screened for biallelic markers. A biallelic marker obtained using this method may show a lower degree of informativeness for conducting association studies, e.g. if the frequency of its less frequent allele may be less than about 10%. Such a biallelic marker will however be sufficiently informative to conduct association studies and it will further be appreciated that including less informative biallelic markers in the genetic analysis studies of the present invention, may allow in some cases the direct identification of causal mutations, which may, depending on their penetrance, be rare mutations.

The following is a description of the various parameters of a preferred method used by the inventors for the identification of the biallelic markers of the present invention.

#### 10 II.A. Genomic DNA Samples

The genomic DNA samples from which the biallelic markers of the present invention are generated are preferably obtained from unrelated individuals corresponding to a heterogeneous population of known ethnic background. The number of individuals from whom DNA samples are obtained can vary substantially, preferably from about 10 to about 15 1000, more preferably from about 50 to about 200 individuals. Usually, DNA samples are collected from at least about 100 individuals in order to have sufficient polymorphic diversity in a given population to identify as many markers as possible and to generate statistically significant results.

As for the source of the genomic DNA to be subjected to analysis, any test sample 20 can be foreseen without any particular limitation. These test samples include biological samples, which can be tested by the methods of the present invention described herein, and include human and animal body fluids such as whole blood, serum, plasma, cerebrospinal fluid, urine, lymph fluids, and various external secretions of the respiratory, intestinal and genitourinary tracts, tears, saliva, milk, white blood cells, myelomas and the like; biological 25 fluids such as cell culture supernatants; fixed tissue specimens including tumor and non-tumor tissue and lymph node tissues; bone marrow aspirates and fixed cell specimens. The preferred source of genomic DNA used in the present invention is from peripheral venous blood of each donor. Techniques to prepare genomic DNA from biological samples are well known to the skilled technician. Details of a preferred embodiment are provided in 30 Example 1. The person skilled in the art can choose to amplify pooled or unpooled DNA samples.

#### II.B. DNA Amplification

The identification of biallelic markers in a sample of genomic DNA may be facilitated through the use of DNA amplification methods. DNA samples can be pooled or

unpooled for the amplification step. DNA amplification techniques are well known to those skilled in the art. Various methods to amplify DNA fragments carrying biallelic markers are further described hereinafter in III.B. The PCR technology is the preferred amplification technique used to identify new biallelic markers.

- 5 In a first embodiment, biallelic markers are identified using genomic sequence information generated by the inventors. Genomic DNA fragments, such as the inserts of the BAC clones described above, are sequenced and used to design primers for the amplification of 500 bp fragments. These 500 bp fragments are amplified from genomic DNA and are scanned for biallelic markers. Primers may be designed using the OSP  
10 software (Hillier L. and Green P., 1991). All primers may contain, upstream of the specific target bases, a common oligonucleotide tail that serves as a sequencing primer. Those skilled in the art are familiar with primer extensions, which can be used for these purposes.

In another embodiment of the invention, genomic sequences of candidate genes are available in public databases allowing direct screening for biallelic markers. Preferred  
15 primers, useful for the amplification of genomic sequences encoding the candidate genes, focus on promoters, exons and splice sites of the genes. A biallelic marker present in these functional regions of the gene have a higher probability to be a causal mutation.

Preferred primers include those disclosed in Figure 7.

## **II.C. Sequencing of Amplified Genomic DNA and Identification of Single Nucleotide**

### **20 Polymorphisms**

The amplification products generated as described above, are then sequenced using any method known and available to the skilled technician. Methods for sequencing DNA using either the dideoxy-mediated method (Sanger method) or the Maxam-Gilbert method are widely known to those of ordinary skill in the art. Such methods are for example  
25 disclosed in Maniatis et al. (Molecular Cloning, A Laboratory Manual, Cold Spring Harbor Press, Second Edition, 1989). Alternative approaches include hybridization to high-density DNA probe arrays as described in Chee et al. (*Science* 274, 610, 1996).

Preferably, the amplified DNA is subjected to automated dideoxy terminator sequencing reactions using a dye-primer cycle sequencing protocol. The products of the  
30 sequencing reactions are run on sequencing gels and the sequences are determined using gel image analysis. The polymorphism search is based on the presence of superimposed peaks in the electrophoresis pattern resulting from different bases occurring at the same position. Because each dideoxy terminator is labeled with a different fluorescent molecule, the two peaks corresponding to a biallelic site present distinct colors corresponding to two different

nucleotides at the same position on the sequence. However, the presence of two peaks can be an artifact due to background noise. To exclude such an artifact, the two DNA strands are sequenced and a comparison between the peaks is carried out. In order to be registered as a polymorphic sequence, the polymorphism has to be detected on both strands.

5       The above procedure permits those amplification products, which contain biallelic markers to be identified. The detection limit for the frequency of biallelic polymorphisms detected by sequencing pools of 100 individuals is approximately 0.1 for the minor allele, as verified by sequencing pools of known allelic frequencies. However, more than 90% of the biallelic polymorphisms detected by the pooling method have a frequency for the minor  
10 allele higher than 0.25. Therefore, the biallelic markers selected by this method have a frequency of at least 0.1 for the minor allele and less than 0.9 for the major allele. Preferably at least 0.2 for the minor allele and less than 0.8 for the major allele, more preferably at least 0.3 for the minor allele and less than 0.7 for the major allele, thus a heterozygosity rate higher than 0.18, preferably higher than 0.32, more preferably higher  
15 than 0.42.

In another embodiment, biallelic markers are detected by sequencing individual DNA samples, the frequency of the minor allele of such a biallelic marker may be less than 0.1.

The markers carried by the same fragment of genomic DNA, such as the insert in a  
20 BAC clone, need not necessarily be ordered with respect to one another within the genomic fragment to conduct association studies. However, in some embodiments of the present invention, the order of biallelic markers carried by the same fragment of genomic DNA are determined.

#### **II.D. Validation of the Biallelic Markers of the Present Invention**

25       The polymorphisms are evaluated for their usefulness as genetic markers by validating that both alleles are present in a population. Validation of the biallelic markers is accomplished by genotyping a group of individuals by a method of the invention and demonstrating that both alleles are present. Microsequencing is a preferred method of genotyping alleles. The validation by genotyping step may be performed on individual  
30 samples derived from each individual in the group or by genotyping a pooled sample derived from more than one individual. The group can be as small as one individual if that individual is heterozygous for the allele in question. Preferably the group contains at least three individuals, more preferably the group contains five or six individuals, so that a single validation test will be more likely to result in the validation of more of the biallelic markers



that are being tested. It should be noted, however, that when the validation test is performed on a small group it may result in a false negative result if as a result of sampling error none of the individuals tested carries one of the two alleles. Thus, the validation process is less useful in demonstrating that a particular initial result is an artifact, than it is at

- 5 demonstrating that there is a *bona fide* biallelic marker at a particular position in a sequence. For an indication of whether a particular biallelic marker has been validated see Figure 1. All of the genotyping, haplotyping, association, and interaction study methods of the invention may optionally be performed solely with validated biallelic markers.

#### **II.E. Evaluation of the Frequency of the Biallelic Markers of the Present Invention**

- 10 The validated biallelic markers are further evaluated for their usefulness as genetic markers by determining the frequency of the least common allele at the biallelic marker site. The determination of the least common allele is accomplished by genotyping a group of individuals by a method of the invention and demonstrating that both alleles are present. This determination of frequency by genotyping step may be performed on individual
- 15 samples derived from each individual in the group or by genotyping a pooled sample derived from more than one individual. The group must be large enough to be representative of the population as a whole. Preferably the group contains at least 20 individuals, more preferably the group contains at least 50 individuals, most preferably the group contains at least 100 individuals. Of course the larger the group the greater the
- 20 accuracy of the frequency determination because of reduced sampling error. For an indication of the frequency for the less common allele of a particular biallelic marker of the invention see Figure 1. A biallelic marker wherein the frequency of the less common allele is 30% or more is termed a "high quality biallelic marker." All of the genotyping, haplotyping, association, and interaction study methods of the invention may optionally be
- 25 performed solely with high quality biallelic markers.

#### **III. Methods of Genotyping an Individual for Biallelic Markers**

- Methods are provided to genotype a biological sample for one or more biallelic markers of the present invention, all of which may be performed *in vitro*. Such methods of genotyping comprise determining the identity of a nucleotide at a DME-related biallelic
- 30 marker by any method known in the art. These methods find use in genotyping case-control populations in association studies as well as individuals in the context of detection of alleles of biallelic markers which, are known to be associated with a given trait, in which case both copies of the biallelic marker present in individual's genome are determined so that an individual may be classified as homozygous or heterozygous for a particular allele.

These genotyping methods can be performed nucleic acid samples derived from a single individual or pooled DNA samples.

Genotyping can be performed using similar methods as those described above for the identification of the biallelic markers, or using other genotyping methods such as those  
5 further described below. In preferred embodiments, the comparison of sequences of amplified genomic fragments from different individuals is used to identify new biallelic markers whereas microsequencing is used for genotyping known biallelic markers in diagnostic and association study applications.

### **III.A. Source of DNA for Genotyping**

10 Any source of nucleic acids, in purified or non-purified form, can be utilized as the starting nucleic acid, provided it contains or is suspected of containing the specific nucleic acid sequence desired. DNA or RNA may be extracted from cells, tissues, body fluids and the like as described above in II.A. "Genomic DNA Samples." While nucleic acids for use in the genotyping methods of the invention can be derived from any mammalian source, the  
15 test subjects and individuals from which nucleic acid samples are taken are generally understood to be human.

### **III.B. Amplification of DNA Fragments Comprising Biallelic Markers**

Methods and polynucleotides are provided to amplify a segment of nucleotides comprising one or more biallelic marker of the present invention. It will be appreciated that  
20 amplification of DNA fragments comprising biallelic markers may be used in various methods and for various purposes and is not restricted to genotyping. Nevertheless, many genotyping methods, although not all, require the previous amplification of the DNA region carrying the biallelic marker of interest. Such methods specifically increase the concentration or total number of sequences that span the biallelic marker or include that site  
25 and sequences located either distal or proximal to it. Diagnostic assays may also rely on amplification of DNA segments carrying a biallelic marker of the present invention.

Amplification of DNA may be achieved by any method known in the art. The established PCR (polymerase chain reaction) method or by developments thereof or alternatives. Amplification methods which can be utilized herein include but are not limited  
30 to Ligase Chain Reaction (LCR) as described in EP A 320 308 and EP A 439 182, Gap LCR (Wolcott, M.J., Clin. Microbiol. Rev. 5:370-386), the so-called "NASBA" or "3SR" technique described in Guatelli J.C. et al. (*Proc. Natl. Acad. Sci. USA* 87:1874-1878, 1990) and in Compton J. (*Nature* 350:91-92, 1991), Q-beta amplification as described in European Patent Application no 4544610, strand displacement amplification as described in

Walker et al. (*Clin. Chem.* 42:9-13, 1996) and EP A 684 315 and, target mediated amplification as described in PCT Publication WO 9322461.

LCR and Gap LCR are exponential amplification techniques, both depend on DNA ligase to join adjacent primers annealed to a DNA molecule. In Ligase Chain Reaction (LCR), probe pairs are used which include two primary (first and second) and two secondary (third and fourth) probes, all of which are employed in molar excess to target. The first probe hybridizes to a first segment of the target strand and the second probe hybridizes to a second segment of the target strand, the first and second segments being contiguous so that the primary probes abut one another in 5' phosphate-3'hydroxyl relationship, and so that a ligase can covalently fuse or ligate the two probes into a fused product. In addition, a third (secondary) probe can hybridize to a portion of the first probe and a fourth (secondary) probe can hybridize to a portion of the second probe in a similar abutting fashion. Of course, if the target is initially double stranded, the secondary probes also will hybridize to the target complement in the first instance. Once the ligated strand of primary probes is separated from the target strand, it will hybridize with the third and fourth probes which can be ligated to form a complementary, secondary ligated product. It is important to realize that the ligated products are functionally equivalent to either the target or its complement. By repeated cycles of hybridization and ligation, amplification of the target sequence is achieved. A method for multiplex LCR has also been described (WO 9320227). Gap LCR (GLCR) is a version of LCR where the probes are not adjacent but are separated by 2 to 3 bases.

For amplification of mRNAs, it is within the scope of the present invention to reverse transcribe mRNA into cDNA followed by polymerase chain reaction (RT-PCR); or, to use a single enzyme for both steps as described in U.S. Patent No. 5,322,770 or, to use Asymmetric Gap LCR (RT-AGLCR) as described by Marshall R.L. et al. (*PCR Methods and Applications* 4:80-84, 1994). AGLCR is a modification of GLCR that allows the amplification of RNA.

Some of these amplification methods are particularly suited for the detection of single nucleotide polymorphisms and allow the simultaneous amplification of a target sequence and the identification of the polymorphic nucleotide as it is further described in IIIC.

The PCR technology is the preferred amplification technique used in the present invention. A variety of PCR techniques are familiar to those skilled in the art. For a review of PCR technology, see Molecular Cloning to Genetic Engineering White, B.A. Ed. in *Methods*

in *Molecular Biology* 67: Humana Press, Totowa (1997) and the publication entitled "PCR Methods and Applications" (1991, Cold Spring Harbor Laboratory Press). In each of these PCR procedures, PCR primers on either side of the nucleic acid sequences to be amplified are added to a suitably prepared nucleic acid sample along with dNTPs and a thermostable  
5 polymerase such as Taq polymerase, Pfu polymerase, or Vent polymerase. The nucleic acid in the sample is denatured and the PCR primers are specifically hybridized to complementary nucleic acid sequences in the sample. The hybridized primers are extended. Thereafter, another cycle of denaturation, hybridization, and extension is initiated. The cycles are repeated multiple times to produce an amplified fragment containing the nucleic acid sequence between  
10 the primer sites. PCR has further been described in several patents including US Patents 4,683,195, 4,683,202 and 4,965,188.

The identification of biallelic markers as described above allows the design of appropriate oligonucleotides, which can be used as primers to amplify DNA fragments comprising the biallelic markers of the present invention. Amplification can be performed  
15 using the primers initially used to discover new biallelic markers which are described herein or any set of primers allowing the amplification of a DNA fragment comprising a biallelic marker of the present invention. Primers can be prepared by any suitable method. As for example, direct chemical synthesis by a method such as the phosphodiester method of Narang S.A. et al. (*Methods Enzymol.* 68:90-98, 1979), the phosphodiester method of  
20 Brown E.L. et al. (*Methods Enzymol.* 68:109-151, 1979), the diethylphosphoramidite method of Beaucage et al. (*Tetrahedron Lett.* 22:1859-1862, 1981) and the solid support method described in EP 0 707 592.

In some embodiments the present invention provides primers for amplifying a DNA fragment containing one or more biallelic markers of the present invention. Preferred  
25 amplification primers are listed in Figure 7. It will be appreciated that the primers listed are merely exemplary and that any other set of primers which produce amplification products containing one or more biallelic markers of the present invention.

The primers are selected to be substantially complementary to the different strands of each specific sequence to be amplified. The length of the primers of the present  
30 invention can range from 8 to 100 nucleotides, preferably from 8 to 50, 8 to 30 or more preferably 8 to 25 nucleotides. Shorter primers tend to lack specificity for a target nucleic acid sequence and generally require cooler temperatures to form sufficiently stable hybrid complexes with the template. Longer primers are expensive to produce and can sometimes self-hybridize to form hairpin structures. The formation of stable hybrids depends on the

melting temperature ( $T_m$ ) of the DNA. The  $T_m$  depends on the length of the primer, the ionic strength of the solution and the G+C content. The higher the G+C content of the primer, the higher is the melting temperature because G:C pairs are held by three H bonds whereas A:T pairs have only two. The G+C content of the amplification primers of the  
5 present invention preferably ranges between 10 and 75 %, more preferably between 35 and 60 %, and most preferably between 40 and 55 %. The appropriate length for primers under a particular set of assay conditions may be empirically determined by one of skill in the art.

The spacing of the primers determines the length of the segment to be amplified. In the context of the present invention amplified segments carrying biallelic markers can range  
10 in size from at least about 25 bp to 35 kbp. Amplification fragments from 25-3000 bp are typical, fragments from 50-1000 bp are preferred and fragments from 100-600 bp are highly preferred. It will be appreciated that amplification primers for the biallelic markers may be any sequence which allow the specific amplification of any DNA fragment carrying the markers. Amplification primers may be labeled or immobilized on a solid support as  
15 described in I.

### III.C. Methods of Genotyping DNA samples for Biallelic Markers

Any method known in the art can be used to identify the nucleotide present at a biallelic marker site. Since the biallelic marker allele to be detected has been identified and specified in the present invention, detection will prove simple for one of ordinary skill in the  
20 art by employing any of a number of techniques. Many genotyping methods require the previous amplification of the DNA region carrying the biallelic marker of interest. While the amplification of target or signal is often preferred at present, ultrasensitive detection methods which do not require amplification are also encompassed by the present genotyping methods. Methods well-known to those skilled in the art that can be used to detect biallelic  
25 polymorphisms include methods such as, conventional dot blot analyzes, single strand conformational polymorphism analysis (SSCP) described by Orita et al. (*Proc. Natl. Acad. Sci. U.S.A* 86:27776-2770, 1989), denaturing gradient gel electrophoresis (DGGE), heteroduplex analysis, mismatch cleavage detection, and other conventional techniques as described in Sheffield, V.C. et al. (*Proc. Natl. Acad. Sci. USA* 49:699-706, 1991), White et  
30 al. (*Genomics* 12:301-306, 1992), Grompe, M. et al. (*Proc. Natl. Acad. Sci. USA* 86:5855-5892, 1989) and Grompe, M. (*Nature Genetics* 5:111-117, 1993). Another method for determining the identity of the nucleotide present at a particular polymorphic site employs a specialized exonuclease-resistant nucleotide derivative as described in US patent 4,656,127.

Preferred methods involve directly determining the identity of the nucleotide present at a biallelic marker site by sequencing assay, enzyme-based mismatch detection assay, or hybridization assay. The following is a description of some preferred methods. A highly preferred method is the microsequencing technique. The term "sequencing assay" is used  
5 herein to refer to polymerase extension of duplex primer/template complexes and includes both traditional sequencing and microsequencing.

### 1) Sequencing assays

The nucleotide present at a polymorphic site can be determined by sequencing methods. In a preferred embodiment, DNA samples are subjected to PCR amplification  
10 before sequencing as described above. DNA sequencing methods are described in IIC.

Preferably, the amplified DNA is subjected to automated dideoxy terminator sequencing reactions using a dye-primer cycle sequencing protocol. Sequence analysis allows the identification of the base present at the biallelic marker site.

### 2) Microsequencing assays

15 In microsequencing methods, a nucleotide at the polymorphic site that is unique to one of the alleles in a target DNA is detected by a single nucleotide primer extension reaction. This method involves appropriate microsequencing primers which, hybridize just upstream of a polymorphic base of interest in the target nucleic acid. A polymerase is used to specifically extend the 3' end of the primer with one single ddNTP (chain terminator)  
20 complementary to the selected nucleotide at the polymorphic site. Next the identity of the incorporated nucleotide is determined in any suitable way.

Typically, microsequencing reactions are carried out using fluorescent ddNTPs and the extended microsequencing primers are analyzed by electrophoresis on ABI 377 sequencing machines to determine the identity of the incorporated nucleotide as described in  
25 EP 412 883. Alternatively capillary electrophoresis can be used in order to process a higher number of assays simultaneously. An example of a typical microsequencing procedure that can be used in the context of the present invention is provided in Example 2.

Different approaches can be used to detect the nucleotide added to the microsequencing primer. A homogeneous phase detection method based on fluorescence  
30 resonance energy transfer has been described by Chen and Kwok (*Nucleic Acids Research* 25:347-353 1997) and Chen et al. (*Proc. Natl. Acad. Sci. USA* 94/20 10756-10761, 1997). In this method amplified genomic DNA fragments containing polymorphic sites are incubated with a 5'-fluorescein-labeled primer in the presence of allelic dye-labeled dideoxyribonucleoside triphosphates and a modified Taq polymerase. The dye-labeled

primer is extended one base by the dye-terminator specific for the allele present on the template. At the end of the genotyping reaction, the fluorescence intensities of the two dyes in the reaction mixture are analyzed directly without separation or purification. All these steps can be performed in the same tube and the fluorescence changes can be monitored in  
5 real time. Alternatively, the extended primer may be analyzed by MALDI-TOF Mass Spectrometry. The base at the polymorphic site is identified by the mass added onto the microsequencing primer (see Haff L.A. and Smirnov I.P., *Genome Research*, 7:378-388, 1997).

Microsequencing may be achieved by the established microsequencing method or by  
10 developments or derivatives thereof. Alternative methods include several solid-phase microsequencing techniques. The basic microsequencing protocol is the same as described previously, except that the method is conducted as a heterogenous phase assay, in which the primer or the target molecule is immobilized or captured onto a solid support. To simplify the primer separation and the terminal nucleotide addition analysis, oligonucleotides are  
15 attached to solid supports or are modified in such ways that permit affinity separation as well as polymerase extension. The 5' ends and internal nucleotides of synthetic oligonucleotides can be modified in a number of different ways to permit different affinity separation approaches, e.g., biotinylation. If a single affinity group is used on the oligonucleotides, the oligonucleotides can be separated from the incorporated terminator  
20 reagent. This eliminates the need of physical or size separation. More than one oligonucleotide can be separated from the terminator reagent and analyzed simultaneously if more than one affinity group is used. This permits the analysis of several nucleic acid species or more nucleic acid sequence information per extension reaction. The affinity group need not be on the priming oligonucleotide but could alternatively be present on the  
25 template. For example, immobilization can be carried out via an interaction between biotinylated DNA and streptavidin-coated microtitration wells or avidin-coated polystyrene particles. In the same manner oligonucleotides or templates may be attached to a solid support in a high-density format. In such solid phase microsequencing reactions, incorporated ddNTPs can be radiolabeled (Syvänen, *Clinica Chimica Acta* 226:225-236,  
30 1994) or linked to fluorescein (Livak and Hainer, *Human Mutation* 3:379-385, 1994). The detection of radiolabeled ddNTPs can be achieved through scintillation-based techniques. The detection of fluorescein-linked ddNTPs can be based on the binding of anti fluorescein antibody conjugated with alkaline phosphatase, followed by incubation with a chromogenic substrate (such as *p*-nitrophenyl phosphate). Other possible reporter-detection pairs include:

ddNTP linked to dinitrophenyl (DNP) and anti-DNP alkaline phosphatase conjugate (Harju et al., *Clin. Chem.* 39/11 2282-2287, 1993) or biotinylated ddNTP and horseradish peroxidase-conjugated streptavidin with *o*-phenylenediamine as a substrate (WO 92/15712). As yet another alternative solid-phase microsequencing procedure, Nyren et al. (*Analytical Biochemistry* 208:171-175, 1993) described a method relying on the detection of DNA polymerase activity by an enzymatic luminometric inorganic pyrophosphate detection assay (ELIDA).

Pastinen et al. (*Genome research* 7:606-614, 1997), describe a method for multiplex detection of single nucleotide polymorphism in which the solid phase minisequencing principle is applied to an oligonucleotide array format. High-density arrays of DNA probes attached to a solid support (DNA chips) are further described in III.C.5.

In one aspect the present invention provides polynucleotides and methods to genotype one or more biallelic markers of the present invention by performing a microsequencing assay. Preferred microsequencing primers include those being featured Figure 6. It will be appreciated that the microsequencing primers listed in Figure 6 are merely exemplary and that, any primer having a 3' end immediately adjacent to a polymorphic nucleotide may be used. Similarly, it will be appreciated that microsequencing analysis may be performed for any biallelic marker or any combination of biallelic markers of the present invention. One aspect of the present invention is a solid support which includes one or more microsequencing primers listed in Figure 6, or fragments comprising at least 8, at least 12, at least 15, or at least 20 consecutive nucleotides thereof and having a 3' terminus immediately upstream of the corresponding biallelic marker, for determining the identity of a nucleotide at biallelic marker site.

### **3) Mismatch detection assays based on polymerases and ligases**

In one aspect the present invention provides polynucleotides and methods to determine the allele of one or more biallelic markers of the present invention in a biological sample, by mismatch detection assays based on polymerases and/or ligases. These assays are based on the specificity of polymerases and ligases. Polymerization reactions places particularly stringent requirements on correct base pairing of the 3' end of the amplification primer and the joining of two oligonucleotides hybridized to a target DNA sequence is quite sensitive to mismatches close to the ligation site, especially at the 3' end. The terms "enzyme based mismatch detection assay" are used herein to refer to any method of determining the allele of a biallelic marker based on the specificity of ligases and polymerases. Preferred methods are described below. Methods, primers and various



parameters to amplify DNA fragments comprising biallelic markers of the present invention are further described above in III.B.

#### **Allele specific amplification**

Discrimination between the two alleles of a biallelic marker can also be achieved by  
5 allele specific amplification, a selective strategy, whereby one of the alleles is amplified without amplification of the other allele. This is accomplished by placing a polymorphic base at the 3' end of one of the amplification primers. Because the extension forms from the 3' end of the primer, a mismatch at or near this position has an inhibitory effect on  
10 amplification. Therefore, under appropriate amplification conditions, these primers only direct amplification on their complementary allele. Designing the appropriate allele-specific primer and the corresponding assay conditions are well with the ordinary skill in the art.

#### **Ligation/amplification based methods**

The "Oligonucleotide Ligation Assay" (OLA) uses two oligonucleotides which are designed to be capable of hybridizing to abutting sequences of a single strand of a target  
15 molecules. One of the oligonucleotides is biotinylated, and the other is detectably labeled. If the precise complementary sequence is found in a target molecule, the oligonucleotides will hybridize such that their termini abut, and create a ligation substrate that can be captured and detected. OLA is capable of detecting biallelic markers and may be advantageously combined with PCR as described by Nickerson D.A. et al. (*Proc. Natl. Acad. Sci. U.S.A.*  
20 87:8923-8927, 1990). In this method, PCR is used to achieve the exponential amplification of target DNA, which is then detected using OLA.

Other methods which are particularly suited for the detection of biallelic markers include LCR (ligase chain reaction), Gap LCR (GLCR) which are described above in III.B. As mentioned above LCR uses two pairs of probes to exponentially amplify a specific  
25 target. The sequences of each pair of oligonucleotides, is selected to permit the pair to hybridize to abutting sequences of the same strand of the target. Such hybridization forms a substrate for a template-dependant ligase. In accordance with the present invention, LCR can be performed with oligonucleotides having the proximal and distal sequences of the same strand of a biallelic marker site. In one embodiment, either oligonucleotide will be  
30 designed to include the biallelic marker site. In such an embodiment, the reaction conditions are selected such that the oligonucleotides can be ligated together only if the target molecule either contains or lacks the specific nucleotide(s) that is complementary to the biallelic marker on the oligonucleotide. In an alternative embodiment, the oligonucleotides will not include the biallelic marker, such that when they hybridize to the target molecule, a "gap" is

created as described in WO 90/01069. This gap is then "filled" with complementary dNTPs (as mediated by DNA polymerase), or by an additional pair of oligonucleotides. Thus at the end of each cycle, each single strand has a complement capable of serving as a target during the next cycle and exponential allele-specific amplification of the desired sequence is  
5 obtained.

Ligase/Polymerase-mediated Genetic Bit Analysis<sup>TM</sup> is another method for determining the identity of a nucleotide at a preselected site in a nucleic acid molecule (WO 95/21271). This method involves the incorporation of a nucleoside triphosphate that is complementary to the nucleotide present at the preselected site onto the terminus of a primer  
10 molecule, and their subsequent ligation to a second oligonucleotide. The reaction is monitored by detecting a specific label attached to the reaction's solid phase or by detection in solution.

#### 4) Hybridization assay methods

A preferred method of determining the identity of the nucleotide present at a biallelic  
15 marker site involves nucleic acid hybridization. The hybridization probes, which can be conveniently used in such reactions, preferably include the probes defined herein. Any hybridization assay may be used including Southern hybridization, Northern hybridization, dot blot hybridization and solid-phase hybridization (see Sambrook et al., Molecular Cloning – A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., 1989).

20 Hybridization refers to the formation of a duplex structure by two single stranded nucleic acids due to complementary base pairing. Hybridization can occur between exactly complementary nucleic acid strands or between nucleic acid strands that contain minor regions of mismatch. Specific probes can be designed that hybridize to one form of a biallelic marker and not to the other and therefore are able to discriminate between different  
25 allelic forms. Allele-specific probes are often used in pairs, one member of a pair showing perfect match to a target sequence containing the original allele and the other showing a perfect match to the target sequence containing the alternative allele. Hybridization conditions should be sufficiently stringent that there is a significant difference in hybridization intensity between alleles, and preferably an essentially binary response,  
30 whereby a probe hybridizes to only one of the alleles. Stringent, sequence specific hybridization conditions, under which a probe will hybridize only to the exactly complementary target sequence are well known in the art (Sambrook et al., Molecular Cloning – A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., 1989). Stringent conditions are sequence dependent and will be different in different

circumstances. Generally, stringent conditions are selected to be about 5°C lower than the thermal melting point (T<sub>m</sub>) for the specific sequence at a defined ionic strength and pH. By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 5 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 10<sup>6</sup> cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the 10 presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute 15 intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the 20 hybridized probes are detectable by autoradiography. Other conditions of high and intermediate stringency which may be used are well known in the art and as cited in Sambrook et al. (Molecular Cloning - A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., 1989) and Ausubel et al. (Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y., 1989).

25 Although such hybridizations can be performed in solution, it is preferred to employ a solid-phase hybridization assay. The target DNA comprising a biallelic marker of the present invention may be amplified prior to the hybridization reaction. The presence of a specific allele in the sample is determined by detecting the presence or the absence of stable hybrid duplexes formed between the probe and the target DNA. The detection of hybrid 30 duplexes can be carried out by a number of methods. Various detection assay formats are well known which utilize detectable labels bound to either the target or the probe to enable detection of the hybrid duplexes. Typically, hybridization duplexes are separated from unhybridized nucleic acids and the labels bound to the duplexes are then detected. Those skilled in the art will recognize that wash steps may be employed to wash away excess

target DNA or probe. Standard heterogeneous assay formats are suitable for detecting the hybrids using the labels present on the primers and probes.

Two recently developed assays allow hybridization-based allele discrimination with no need for separations or washes (see Landegren U. et al., *Genome Research*, 8:769-776, 1998). The TaqMan assay takes advantage of the 5' nuclease activity of Taq DNA polymerase to digest a DNA probe annealed specifically to the accumulating amplification product. TaqMan probes are labeled with a donor-acceptor dye pair that interacts via fluorescence energy transfer. Cleavage of the TaqMan probe by the advancing polymerase during amplification dissociates the donor dye from the quenching acceptor dye, greatly increasing the donor fluorescence. All reagents necessary to detect two allelic variants can be assembled at the beginning of the reaction and the results are monitored in real time (see Livak et al., *Nature Genetics*, 9:341-342, 1995). In an alternative homogeneous hybridization-based procedure, molecular beacons are used for allele discriminations. Molecular beacons are hairpin-shaped oligonucleotide probes that report the presence of specific nucleic acids in homogeneous solutions. When they bind to their targets they undergo a conformational reorganization that restores the fluorescence of an internally quenched fluorophore (Tyagi et al., *Nature Biotechnology*, 16:49-53, 1998).

The polynucleotides provided herein can be used in hybridization assays for the detection of biallelic marker alleles in biological samples. These probes are characterized in that they preferably comprise between 8 and 50 nucleotides, and in that they are sufficiently complementary to a sequence comprising a biallelic marker of the present invention to hybridize thereto and preferably sufficiently specific to be able to discriminate the targeted sequence for only one nucleotide variation. The GC content in the probes of the invention usually ranges between 10 and 75 %, preferably between 35 and 60 %, and more preferably between 40 and 55 %. The length of these probes can range from 10, 15, 20, or 30 to at least 100 nucleotides, preferably from 10 to 50, more preferably from 18 to 35 nucleotides. A particularly preferred probe is 25 nucleotides in length. Preferably the biallelic marker is within 4 nucleotides of the center of the polynucleotide probe. In particularly preferred probes the biallelic marker is at the center of said polynucleotide. Shorter probes may lack specificity for a target nucleic acid sequence and generally require cooler temperatures to form sufficiently stable hybrid complexes with the template. Longer probes are expensive to produce and can sometimes self-hybridize to form hairpin structures. Methods for the synthesis of oligonucleotide probes have been described above and can be applied to the probes of the present invention.

Preferably the probes of the present invention are labeled or immobilized on a solid support. Labels and solid supports are further described in I. Detection probes are generally nucleic acid sequences or uncharged nucleic acid analogs such as, for example peptide nucleic acids which are disclosed in International Patent Application WO 92/20702, d,  
5 morpholino analogs which are described in U.S. Patents Numbered 5,185,444; 5,034,506 and 5,142,047. The probe may have to be rendered "non-extendable" in that additional dNTPs cannot be added to the probe. In and of themselves analogs usually are non-extendable and nucleic acid probes can be rendered non-extendable by modifying the 3' end of the probe such that the hydroxyl group is no longer capable of participating in elongation.  
10 For example, the 3' end of the probe can be functionalized with the capture or detection label to thereby consume or otherwise block the hydroxyl group. Alternatively, the 3' hydroxyl group simply can be cleaved, replaced or modified, U.S. Patent Application Serial No. 07/049,061 filed April 19, 1993 describes modifications, which can be used to render a probe non-extendable.

15 The probes of the present invention are useful for a number of purposes. They can be used in Southern hybridization to genomic DNA or Northern hybridization to mRNA. The probes can also be used to detect PCR amplification products. By assaying the hybridization to an allele specific probe, one can detect the presence or absence of a biallelic marker allele in a given sample.

20 High-Throughput parallel hybridizations in array format are specifically encompassed within "hybridization assays" and are described below.

#### **Hybridization to addressable arrays of oligonucleotides**

Hybridization assays based on oligonucleotide arrays rely on the differences in hybridization stability of short oligonucleotides to perfectly matched and mismatched target  
25 sequence variants. Efficient access to polymorphism information is obtained through a basic structure comprising high-density arrays of oligonucleotide probes attached to a solid support (the chip) at selected positions. Each DNA chip can contain thousands to millions of individual synthetic DNA probes arranged in a grid-like pattern and miniaturized to the size of a dime.

30 The chip technology has already been applied with success in numerous cases. For example, the screening of mutations has been undertaken in the BRCA1 gene, in *S. cerevisiae* mutant strains, and in the protease gene of HIV-1 virus (Hacia et al., *Nature Genetics*, 14(4):441-447, 1996; Shoemaker et al., *Nature Genetics*, 14(4):450-456, 1996 ; Kozal et al., *Nature Medicine*, 2:753-759, 1996). Chips of various formats for use in

detecting biallelic polymorphisms can be produced on a customized basis by Affymetrix (GeneChip™), Hyseq (HyChip and HyGnostics), and Protogene Laboratories.

In general, these methods employ arrays of oligonucleotide probes that are complementary to target nucleic acid sequence segments from an individual which, target  
5 sequences include a polymorphic marker. EP785280, describes a tiling strategy for the detection of single nucleotide polymorphisms. Briefly, arrays may generally be “tilled” for a large number of specific polymorphisms. By “tiling” is generally meant the synthesis of a defined set of oligonucleotide probes which is made up of a sequence complementary to the target sequence of interest, as well as preselected variations of that sequence, e.g.,  
10 substitution of one or more given positions with one or more members of the basis set of monomers, i.e. nucleotides. Tiling strategies are further described in PCT application No. WO 95/11995. In a particular aspect, arrays are tiled for a number of specific, identified biallelic marker sequences. In particular the array is tiled to include a number of detection blocks, each detection block being specific for a specific biallelic marker or a set of biallelic  
15 markers. For example, a detection block may be tiled to include a number of probes, which span the sequence segment that includes a specific polymorphism. To ensure probes that are complementary to each allele, the probes are synthesized in pairs differing at the biallelic marker. In addition to the probes differing at the polymorphic base, monosubstituted probes are also generally tiled within the detection block. These  
20 monosubstituted probes have bases at and up to a certain number of bases in either direction from the polymorphism, substituted with the remaining nucleotides (selected from A, T, G, C and U). Typically the probes in a tiled detection block will include substitutions of the sequence positions up to and including those that are 5 bases away from the biallelic marker. The monosubstituted probes provide internal controls for the tiled array, to  
25 distinguish actual hybridization from artefactual cross-hybridization. Upon completion of hybridization with the target sequence and washing of the array, the array is scanned to determine the position on the array to which the target sequence hybridizes. The hybridization data from the scanned array is then analyzed to identify which allele or alleles of the biallelic marker are present in the sample. Hybridization and scanning may be carried  
30 out as described in PCT application No. WO 92/10092 and WO 95/11995 and US patent No. 5,424,186, .

Thus, in some embodiments, the chips may comprise an array of nucleic acid sequences of fragments of about 15 nucleotides in length. In further embodiments, the chip may comprise an array including at least one of the sequences selected from the group

consisting of SEQ ID No. 1-38, 40-54, 56-463, 465-487, 490-493 and the sequences complementary thereto, or a fragment thereof at least about 8 consecutive nucleotides, preferably 10, 15, 20, more preferably 25, 30, 40, 47, or 50 consecutive nucleotides. In some embodiments, the chip may comprise an array of at least 2, 3, 4, 5, 6, 7, 8 or more of these polynucleotides of the invention. Solid supports and polynucleotides of the present invention attached to solid supports are further described in I. "Biallelic Markers and Polynucleotides Comprising Biallelic Markers."

#### 5) Integrated Systems

Another technique, which may be used to analyze polymorphisms, includes multicomponent integrated systems, which miniaturize and compartmentalize processes such as PCR and capillary electrophoresis reactions in a single functional device. An example of such technique is disclosed in US patent 5,589,136, which describes the integration of PCR amplification and capillary electrophoresis in chips.

Integrated systems can be envisaged mainly when microfluidic systems are used. These systems comprise a pattern of microchannels designed onto a glass, silicon, quartz, or plastic wafer included on a microchip. The movements of the samples are controlled by electric, electroosmotic or hydrostatic forces applied across different areas of the microchip. For genotyping biallelic markers, the microfluidic system may integrate nucleic acid amplification, microsequencing, capillary electrophoresis and a detection method such as laser-induced fluorescence detection.

#### IV. Methods of Genetic Analysis Using the Biallelic Markers of the Present Invention

Different methods are available for the genetic analysis of complex traits (see Lander and Schork, *Science*, 265, 2037-2048, 1994). The search for disease-susceptibility genes is conducted using two main methods: the linkage approach in which evidence is sought for cosegregation between a locus and a putative trait locus using family studies, and the association approach in which evidence is sought for a statistically significant association between an allele and a trait or a trait causing allele (Khoury J. et al., *Fundamentals of Genetic Epidemiology*, Oxford University Press, NY, 1993). In general, the biallelic markers of the present invention find use in any method known in the art to demonstrate a statistically significant correlation between a genotype and a phenotype. The biallelic markers may be used in parametric and non-parametric linkage analysis methods. Preferably, the biallelic markers of the present invention are used to identify genes associated with detectable traits using association studies, an approach which does not

require the use of affected families and which permits the identification of genes associated with complex and sporadic traits.

The genetic analysis using the biallelic markers of the present invention may be conducted on any scale. The whole set of biallelic markers of the present invention or any  
5 subset of biallelic markers of the present invention may be used. In some embodiments a subset of biallelic markers corresponding to one or several candidate genes of the present invention may be used. In other embodiments a subset of biallelic markers corresponding to candidate genes from a given metabolic pathway may be used. Such pathways include glucoronidation and glutathione conjugation. Alternatively, a subset of biallelic markers of  
10 the present invention localised on a specific chromosome segment may be used. Further, any set of genetic markers including a biallelic marker of the present invention may be used. A set of biallelic polymorphisms that, could be used as genetic markers in combination with the biallelic markers of the present invention, has been described in WO 98/20165. As mentioned above, it should be noted that the biallelic markers of the present invention may  
15 be included in any complete or partial genetic map of the human genome. These different uses are specifically contemplated in the present invention and claims.

#### **IV.A. Linkage Analysis**

Linkage analysis is based upon establishing a correlation between the transmission of genetic markers and that of a specific trait throughout generations within a family. Thus,  
20 the aim of linkage analysis is to detect marker loci that show cosegregation with a trait of interest in pedigrees.

##### **Parametric methods**

When data are available from successive generations there is the opportunity to study the degree of linkage between pairs of loci. Estimates of the recombination fraction  
25 enable loci to be ordered and placed onto a genetic map. With loci that are genetic markers, a genetic map can be established, and then the strength of linkage between markers and traits can be calculated and used to indicate the relative positions of markers and genes affecting those traits (Weir, B.S., *Genetic data Analysis II: Methods for Discrete population genetic Data*, Sinauer Assoc., Inc., Sunderland, MA, USA, 1996). The classical method for  
30 linkage analysis is the logarithm of odds (lod) score method (see Morton N.E., *Am.J. Hum. Genet.*, 7:277-318, 1955; Ott J., *Analysis of Human Genetic Linkage*, John Hopkins University Press, Baltimore, 1991). Calculation of lod scores requires specification of the mode of inheritance for the disease (parametric method). Generally, the length of the candidate region identified using linkage analysis is between 2 and 20Mb. Once a candidate



region is identified as described above, analysis of recombinant individuals using additional markers allows further delineation of the candidate region. Linkage analysis studies have generally relied on the use of a maximum of 5,000 microsatellite markers, thus limiting the maximum theoretical attainable resolution of linkage analysis to about 600 kb on average.

- 5        Linkage analysis has been successfully applied to map simple genetic traits that show clear Mendelian inheritance patterns and have a high penetrance (i.e., the ratio between the number of trait positive carriers of allele *a* and the total number of *a* carriers in the population). However, parametric linkage analysis suffers from a variety of drawbacks. First, it is limited by its reliance on the choice of a genetic model suitable for each studied
- 10 trait. Furthermore, as already mentioned, the resolution attainable using linkage analysis is limited, and complementary studies are required to refine the analysis of the typical 2Mb to 20Mb regions initially identified through linkage analysis. In addition, parametric linkage analysis approaches have proven difficult when applied to complex genetic traits, such as those due to the combined action of multiple genes and/or environmental factors. It is very
- 15 difficult to model these factors adequately in a lod score analysis. In such cases, too large an effort and cost are needed to recruit the adequate number of affected families required for applying linkage analysis to these situations, as recently discussed by Risch, N. and Merikangas, K. (*Science*, 273:1516-1517, 1996).

#### **Non-parametric methods**

- 20        The advantage of the so-called non-parametric methods for linkage analysis is that they do not require specification of the mode of inheritance for the disease, they tend to be more useful for the analysis of complex traits. In non-parametric methods, one tries to prove that the inheritance pattern of a chromosomal region is not consistent with random Mendelian segregation by showing that affected relatives inherit identical copies of the
- 25 region more often than expected by chance. Affected relatives should show excess "allele sharing" even in the presence of incomplete penetrance and polygenic inheritance. In non-parametric linkage analysis the degree of agreement at a marker locus in two individuals can be measured either by the number of alleles identical by state (IBS) or by the number of alleles identical by descent (IBD). Affected sib pair analysis is a well-known special case
- 30 and is the simplest form of these methods.

The biallelic markers of the present invention may be used in both parametric and non-parametric linkage analysis. Preferably biallelic markers may be used in non-parametric methods which allow the mapping of genes involved in complex traits. The biallelic markers of the present invention may be used in both IBD- and IBS- methods to

map genes affecting a complex trait. In such studies, taking advantage of the high density of biallelic markers, several adjacent biallelic marker loci may be pooled to achieve the efficiency attained by multi-allelic markers (Zhao et al., *Am. J. Hum. Genet.*, 63:225-240, 1998).

- 5        However, both parametric and non-parametric linkage analysis methods analyse affected relatives, they tend to be of limited value in the genetic analysis of drug responses or in the analysis of side effects to treatments. This type of analysis is impractical in such cases due to the lack of availability of familial cases. In fact, the likelihood of having more than one individual in a family being exposed to the same drug at the same time is  
10 extremely low.

#### **IV.B. Population Association Studies**

- The present invention comprises methods for identifying one or several genes among a set of candidate genes that are associated with a detectable trait using the biallelic markers of the present invention. In one embodiment the present invention comprises methods to  
15 detect an association between a biallelic marker allele or a biallelic marker haplotype and a trait. Further, the invention comprises methods to identify a trait causing allele in linkage disequilibrium with any biallelic marker allele of the present invention.

- As described above, alternative approaches can be employed to perform association studies: genome-wide association studies, candidate region association studies and  
20 candidate gene association studies. In a preferred embodiment, the biallelic markers of the present invention are used to perform candidate gene association studies. The candidate gene analysis clearly provides a short-cut approach to the identification of genes and gene polymorphisms related to a particular trait when some information concerning the biology of the trait is available. Further, the biallelic markers of the present invention may be  
25 incorporated in any map of genetic markers of the human genome in order to perform genome-wide association studies. Methods to generate a high-density map of biallelic markers has been described in US Provisional Patent application serial number 60/082,614. The biallelic markers of the present invention may further be incorporated in any map of a specific candidate region of the genome (a specific chromosome or a specific chromosomal  
30 segment for example).

As mentioned above, association studies may be conducted within the general population and are not limited to studies performed on related individuals in affected families. Association studies are extremely valuable as they permit the analysis of sporadic or multifactor traits. Moreover, association studies represent a powerful method for fine-

scale mapping enabling much finer mapping of trait causing alleles than linkage studies.

Studies based on pedigrees often only narrow the location of the trait causing allele.

Association studies using the biallelic markers of the present invention can therefore be used to refine the location of a trait causing allele in a candidate region identified by Linkage

- 5 Analysis methods. Moreover, once a chromosome segment of interest has been identified, the presence of a candidate gene such as a candidate gene of the present invention, in the region of interest can provide a shortcut to the identification of the trait causing allele.

Biallelic markers of the present invention can be used to demonstrate that a candidate gene is associated with a trait. Such uses are specifically contemplated in the present invention

10 and claims.

**1) Determining the frequency of a biallelic marker allele or of a biallelic marker haplotype in a population**

Association studies explore the relationships among frequencies for sets of alleles between loci.

**15 Determining the frequency of an allele in a population**

Allelic frequencies of the biallelic markers in a population can be determined using one of the methods described above under the heading "Methods for genotyping an individual for biallelic markers", or any genotyping procedure suitable for this intended purpose. Genotyping pooled samples or individual samples can determine the frequency of

- 20 a biallelic marker allele in a population. One way to reduce the number of genotypings required is to use pooled samples. A major obstacle in using pooled samples is in terms of accuracy and reproducibility for determining accurate DNA concentrations in setting up the pools. Genotyping individual samples provides higher sensitivity, reproducibility and accuracy and; is the preferred method used in the present invention. Preferably, each
- 25 individual is genotyped separately and simple gene counting is applied to determine the frequency of an allele of a biallelic marker or of a genotype in a given population.

**Determining the frequency of a haplotype in a population**

The gametic phase of haplotypes is unknown when diploid individuals are heterozygous at more than one locus. Using genealogical information in families gametic

- 30 phase can sometimes be inferred (Perlin et al., *Am. J. Hum. Genet.*, 55:777-787, 1994).

When no genealogical information is available different strategies may be used. One possibility is that the multiple-site heterozygous diploids can be eliminated from the analysis, keeping only the homozygotes and the single-site heterozygote individuals, but this approach might lead to a possible bias in the sample composition and the underestimation of

low-frequency haplotypes. Another possibility is that single chromosomes can be studied independently, for example, by asymmetric PCR amplification (see Newton et al., *Nucleic Acids Res.*, 17:2503-2516, 1989; Wu et al., *Proc. Natl. Acad. Sci. USA*, 86:2757, 1989), or by isolation of single chromosome by limit dilution followed by PCR amplification (see Ruano et al., *Proc. Natl. Acad. Sci. USA*, 87:6296-6300, 1990). Further, a sample may be haplotyped for sufficiently close biallelic markers by double PCR amplification of specific alleles (Sarkar, G. and Sommer S.S., *Biotechniques*, 1991). These approaches are not entirely satisfying either because of their technical complexity, the additional cost they entail, their lack of generalisation at a large scale, or the possible biases they introduce. To overcome these difficulties, an algorithm to infer the phase of PCR-amplified DNA genotypes introduced by Clark A.G. (*Mol. Biol. Evol.*, 7:111-122, 1990), may be used. Briefly, the principle is to start filling a preliminary list of haplotypes present in the sample by examining unambiguous individuals, that is, the complete homozygotes and the single-site heterozygotes. Then other individuals in the same sample are screened for the possible occurrence of previously recognised haplotypes. For each positive identification, the complementary haplotype is added to the list of recognised haplotypes, until the phase information for all individuals is either resolved or identified as unresolved. This method assigns a single haplotype to each multiheterozygous individual, whereas several haplotypes are possible when there are more than one heterozygous site. Alternatively, one can use methods estimating haplotype frequencies in a population without assigning haplotypes to each individual. Preferably, a method based on an expectation-maximization (EM) algorithm (Dempster et al., *J. R. Stat. Soc.*, 39B: 1-38, 1977), leading to maximum-likelihood estimates of haplotype frequencies under the assumption of Hardy-Weinberg proportions (random mating) is used (see Excoffier L. and Slatkin M., *Mol. Biol. Evol.*, 12(5): 921-927, 1995). The EM algorithm is a generalised iterative maximum-likelihood approach to estimation that is useful when data are ambiguous and/or incomplete. The EM algorithm is used to resolve heterozygotes into haplotypes. Haplotype estimations are further described below under the heading "Statistical methods." Any other method known in the art to determine or to estimate the frequency of a haplotype in a population may also be used.

## 2) Linkage Disequilibrium analysis

Linkage disequilibrium is the non-random association of alleles at two or more loci and represents a powerful tool for mapping genes involved in disease traits (see Ajioka R.S. et al., *Am. J. Hum. Genet.*, 60:1439-1447, 1997). Biallelic markers, because they are

densely spaced in the human genome and can be genotyped in more numerous numbers than other types of genetic markers (such as RFLP or VNTR markers), are particularly useful in genetic analysis based on linkage disequilibrium. The biallelic markers of the present invention may be used in any linkage disequilibrium analysis method known in the art.

- 5 Briefly, when a disease mutation is first introduced into a population (by a new mutation or the immigration of a mutation carrier), it necessarily resides on a single chromosome and thus on a single "background" or "ancestral" haplotype of linked markers. Consequently, there is complete disequilibrium between these markers and the disease mutation: one finds the disease mutation only in the presence of a specific set of marker
- 10 alleles. Through subsequent generations recombinations occur between the disease mutation and these marker polymorphisms, and the disequilibrium gradually dissipates. The pace of this dissipation is a function of the recombination frequency, so the markers closest to the disease gene will manifest higher levels of disequilibrium than those that are further away. When not broken up by recombination, "ancestral" haplotypes and linkage
- 15 disequilibrium between marker alleles at different loci can be tracked not only through pedigrees but also through populations. Linkage disequilibrium is usually seen as an association between one specific allele at one locus and another specific allele at a second locus.

The pattern or curve of disequilibrium between disease and marker loci is expected

20 to exhibit a maximum that occurs at the disease locus. Consequently, the amount of linkage disequilibrium between a disease allele and closely linked genetic markers may yield valuable information regarding the location of the disease gene. For fine-scale mapping of a disease locus, it is useful to have some knowledge of the patterns of linkage disequilibrium that exist between markers in the studied region. As mentioned above the mapping

25 resolution achieved through the analysis of linkage disequilibrium is much higher than that of linkage studies. The high density of biallelic markers combined with linkage disequilibrium analysis provides powerful tools for fine-scale mapping. Different methods to calculate linkage disequilibrium are described below under the heading "Statistical Methods."

### 30 3) Population-based case-control studies of trait-marker associations

As mentioned above, the occurrence of pairs of specific alleles at different loci on the same chromosome is not random and the deviation from random is called linkage disequilibrium. Association studies focus on population frequencies and rely on the phenomenon of linkage disequilibrium. If a specific allele in a given gene is directly

involved in causing a particular trait, its frequency will be statistically increased in an affected (trait positive) population, when compared to the frequency in a trait negative population or in a random control population. As a consequence of the existence of linkage disequilibrium, the frequency of all other alleles present in the haplotype carrying the trait-causing allele will also be increased in trait positive individuals compared to trait negative individuals or random controls. Therefore, association between the trait and any allele (specifically a biallelic marker allele) in linkage disequilibrium with the trait-causing allele will suffice to suggest the presence of a trait-related gene in that particular region. Case-control populations can be genotyped for biallelic markers to identify associations that narrowly locate a trait causing allele. As any marker in linkage disequilibrium with one given marker associated with a trait will be associated with the trait. Linkage disequilibrium allows the relative frequencies in case-control populations of a limited number of genetic polymorphisms (specifically biallelic markers) to be analysed as an alternative to screening all possible functional polymorphisms in order to find trait-causing alleles. Association studies compare the frequency of marker alleles in unrelated case-control populations, and represent powerful tools for the dissection of complex traits.

#### **Case-control populations (inclusion criteria)**

Population-based association studies do not concern familial inheritance but compare the prevalence of a particular genetic marker, or a set of markers, in case-control populations. They are case-control studies based on comparison of unrelated case (affected or trait positive) individuals and unrelated control (unaffected or trait negative or random) individuals. Preferably the control group is composed of unaffected or trait negative individuals. Further, the control group is ethnically matched to the case population. Moreover, the control group is preferably matched to the case-population for the main known confusion factor for the trait under study (for example age-matched for an age-dependent trait). Ideally, individuals in the two samples are paired in such a way that they are expected to differ only in their disease status. In the following "trait positive population", "case population" and "affected population" are used interchangeably.

An important step in the dissection of complex traits using association studies is the choice of case-control populations (see Lander and Schork, *Science*, 265, 2037-2048, 1994). A major step in the choice of case-control populations is the clinical definition of a given trait or phenotype. Any genetic trait may be analysed by the association method proposed here by carefully selecting the individuals to be included in the trait positive and trait negative phenotypic groups. Four criteria are often useful: clinical phenotype, age at onset,

family history and severity. The selection procedure for continuous or quantitative traits (such as blood pressure for example) involves selecting individuals at opposite ends of the phenotype distribution of the trait under study, so as to include in these trait positive and trait negative populations individuals with non-overlapping phenotypes. Preferably, case-  
5 control populations consist of phenotypically homogeneous populations. Trait positive and trait negative populations consist of phenotypically uniform populations of individuals representing each between 1 and 98%, preferably between 1 and 80%, more preferably between 1 and 50%, and more preferably between 1 and 30%, most preferably between 1 and 20% of the total population under study, and selected among individuals exhibiting non-  
10 overlapping phenotypes. The clearer the difference between the two trait phenotypes, the greater the probability of detecting an association with biallelic markers. The selection of those drastically different but relatively uniform phenotypes enables efficient comparisons in association studies and the possible detection of marked differences at the genetic level, provided that the sample sizes of the populations under study are significant enough.

15 In preferred embodiments, a first group of between 50 and 300 trait positive individuals, preferably about 100 individuals, are recruited according to their phenotypes. A similar number of trait negative individuals are included in such studies.

In the present invention, typical examples of inclusion criteria include a disease involving the metabolic conversion of xenobiotics or the evaluation of the response to a  
20 drug or side effects to treatment with drugs.

Suitable examples of association studies using biallelic markers including the biallelic markers of the present invention, are studies involving the following populations:

a case population suffering from a disease involving the metabolic conversion of xenobiotics and a healthy unaffected control population, or

25 a case population treated with therapeutic agents suffering from side-effects resulting from the treatment and a control population treated with the same agents showing no side-effects, or

a case population treated with therapeutic agents showing a beneficial response and a control population treated with same agents showing no beneficial response.

30 In a preferred embodiment, the trait considered was a side-effect upon drug treatment, the study involved two populations derived from a clinical study of the anti-asthmatic drug zileuton. The case population was composed of asthmatic individuals treated with zileuton showing zileuton-associated hepatotoxicity monitored by the serum level of alanine aminotransferase (ALT) and the control population was composed of

asthmatic individuals treated with zileuton and having no increased serum level of ALT.

Inclusion criteria and association between the biallelic markers of the present invention and zileuton-associated hepatotoxicity are further described below and in Example 4.

#### **Association analysis**

5       The general strategy to perform association studies using biallelic markers derived from a region carrying a candidate gene is to scan two groups of individuals (case-control populations) in order to measure and statistically compare the allele frequencies of the biallelic markers of the present invention in both groups.

      If a statistically significant association with a trait is identified for at least one or  
10 more of the analysed biallelic markers, one can assume that: either the associated allele is directly responsible for causing the trait (the associated allele is the trait causing allele), or more likely the associated allele is in linkage disequilibrium with the trait causing allele. The specific characteristics of the associated allele with respect to the candidate gene function usually gives further insight into the relationship between the associated allele and  
15 the trait (causal or in linkage disequilibrium). If the evidence indicates that the associated allele within the candidate gene is most probably not the trait causing allele but is in linkage disequilibrium with the real trait causing allele, then the trait causing allele can be found by sequencing the vicinity of the associated marker.

      Association studies are usually run in two successive steps. In a first phase, the  
20 frequencies of a reduced number of biallelic markers from one or several candidate genes are determined in the trait positive and trait negative populations. In a second phase of the analysis, the identity of the candidate gene and the position of the genetic loci responsible for the given trait is further refined using a higher density of markers from the relevant region. However, if the candidate gene under study is relatively small in length, as it is the  
25 case for many of the candidate genes analysed included in the present invention, a single phase may be sufficient to establish significant associations.

#### **Haplotype analysis**

      As described above, when a chromosome carrying a disease allele first appears in a population as a result of either mutation or migration, the mutant allele necessarily resides  
30 on a chromosome having a set of linked markers: the ancestral haplotype. This haplotype can be tracked through populations and its statistical association with a given trait can be analysed. Complementing single point (allelic) association studies with multi-point association studies also called haplotype studies increases the statistical power of association studies. Thus, a haplotype association study allows one to define the frequency



and the type of the ancestral carrier haplotype. A haplotype analysis is important in that it increases the statistical power of an analysis involving individual markers.

In a first stage of a haplotype frequency analysis, the frequency of the possible haplotypes based on various combinations of the identified biallelic markers of the invention is determined. The haplotype frequency is then compared for distinct populations of trait positive and control individuals. The number of trait positive individuals, which should be, subjected to this analysis to obtain statistically significant results usually ranges between 30 and 300, with a preferred number of individuals ranging between 50 and 150. The same considerations apply to the number of unaffected individuals (or random control) used in the study. The results of this first analysis provide haplotype frequencies in case-control populations, for each evaluated haplotype frequency a p-value and an odd ratio are calculated. If a statistically significant association is found, the relative risk for an individual carrying the given haplotype of being affected with the trait under study can be approximated.

The preferred 2, 3 and 4 marker haplotypes of the invention are listed in Table 3 below:

**Table 3**

GENE	MARKER 1	MARKER 2	MARKER 3	MARKER 4
MGST-II	12-455-326	12-453-429	12-424-198	
MGST-II	12-455-326	12-453-429	12-424-198	12-454-363
MGST-II	12-447-58	12-455-326	12-461-299	12-453-429
MGST-II	12-441-233	12-461-299	12-453-429	
MGST-II	12-441-233	12-461-299	12-453-429	12-426-154
MGST-II	12-426-154	12-424-198		
MGST-II	12-426-154	12-461-299	12-424-198	
ME1	10-428-219	12-724-225		
UGT1A7	12-128-225	12-156-91	12-139-380	12-140-134
UGT1A7	12-148-311	12-156-91	12-139-380	12-140-134
UGT2B4	10-470-25	12-652-203		
UGT2B4	10-470-25	12-637-219	12-652-203.	

The most preferred 2, 3 and 4 marker haplotypes of the invention are listed in Table 4 below:

**Table 4**

GENE	MARKER 1	MARKER 2	MARKER 3	MARKER 4
MGST-II	12-455-326	12-453-429	12-424-198	
MGST-II	12-455-326	12-453-429	12-424-198	12-454-363
MGST-II	12-447-58	12-455-326	12-461-299	12-453-429
MGST-II	12-426-154	12-424-198		
ME1	10-428-219	12-724-225		

UGT1A7	12-128-225	12-156-91	12-139-380	12-140-134
UGT2B4	10-470-25	12-652-203		

### Interaction Analysis

The biallelic markers of the present invention may also be used to identify patterns of biallelic markers associated with detectable traits resulting from polygenic interactions.

- 5 The analysis of genetic interaction between alleles at unlinked loci requires individual genotyping using the techniques described herein. The analysis of allelic interaction among a selected set of biallelic markers with appropriate level of statistical significance can be considered as a haplotype analysis. Interaction analysis consists in stratifying the case-control populations with respect to a given haplotype for the first loci and performing a
- 10 haplotype analysis with the second loci with each subpopulation.

Statistical methods used in association studies are further described below in IV.C. "Statistical Methods."

#### 4) Testing for linkage in the presence of association

- The biallelic markers of the present invention may further be used in TDT
- 15 (transmission/disequilibrium test). TDT tests for both linkage and association and is not affected by population stratification. TDT requires data for affected individuals and their parents or data from unaffected sibs instead of from parents (see Spielmann S. et al., *Am. J. Hum. Genet.*, 52:506-516, 1993; Schaid D.J. et al., *Genet. Epidemiol.*, 13:423-450, 1996, Spielmann S. and Ewens W.J., *Am. J. Hum. Genet.*, 62:450-458, 1998). Such combined
- 20 tests generally reduce the false-positive errors produced by separate analyses.

### IV.C. Statistical Methods

In general, any method known in the art to test whether a trait and a genotype show a statistically significant correlation may be used.

#### 1) Methods in linkage analysis

- 25 Statistical methods and computer programs useful for linkage analysis are well-known to those skilled in the art (see Terwilliger J.D. and Ott J., *Handbook of Human Genetic Linkage*, John Hopkins University Press, London, 1994; Ott J., *Analysis of Human Genetic Linkage*, John Hopkins University Press, Baltimore, 1991).

#### 2) Methods to estimate haplotype frequencies in a population

- 30 As described above, when genotypes are scored, it is often not possible to distinguish heterozygotes so that haplotype frequencies cannot be easily inferred. When the gametic phase is not known, haplotype frequencies can be estimated from the multilocus genotypic data. Any method known to person skilled in the art can be used to estimate

haplotype frequencies (see Lange K., *Mathematical and Statistical Methods for Genetic Analysis*, Springer, New York, 1997; Weir, B.S., *Genetic data Analysis II: Methods for Discrete population genetic Data*, Sinauer Assoc., Inc., Sunderland, MA, USA, 1996).

Preferably, maximum-likelihood haplotype frequencies are computed using an Expectation-  
 5 Maximization (EM) algorithm (see Dempster et al., *J. R. Stat. Soc.*, 39B:1-38, 1977; Excoffier L. and Slatkin M., *Mol. Biol. Evol.*, 12(5): 921-927, 1995). This procedure is an iterative process aiming at obtaining maximum-likelihood estimates of haplotype frequencies from multi-locus genotype data when the gametic phase is unknown. Haplotype estimations are usually performed by applying the EM algorithm using for example the EM-  
 10 HAPLO program (Hawley M.E. et al., *Am. J. Phys. Anthropol.*, 18:104, 1994) or the Arlequin program (Schneider et al., *Arlequin: a software for population genetics data analysis*, University of Geneva, 1997). The EM algorithm is a generalised iterative maximum likelihood approach to estimation and is briefly described below.

In the following part of this text, phenotypes will refer to multi-locus genotypes with  
 15 unknown phase. Genotypes will refer to known-phase multi-locus genotypes.

Suppose a sample of N unrelated individuals typed for K markers. The data observed are the unknown-phase K-locus phenotypes that can be categorised in F different phenotypes. Suppose that we have H underlying possible haplotypes (in case of K biallelic markers,  $H=2^K$ ).

20 For phenotype j, suppose that  $c_j$  genotypes are possible. We thus have the following equation

$$P_j = \sum_{i=1}^{c_j} pr(genotype_i) = \sum_{i=1}^{c_j} pr(h_k, h_l) \quad \text{Equation 1}$$

where  $P_j$  is the probability of the phenotype j,  $h_k$  and  $h_l$  are the two haplotypes constituent the genotype i. Under the Hardy-Weinberg equilibrium,  $pr(h_k, h_l)$  becomes :

25  $pr(h_k, h_l) = pr(h_k)^2$  if  $h_k = h_l$ ,  $pr(h_k, h_l) = 2pr(h_k).pr(h_l)$  if  $h_k \neq h_l$ . Equation 2

The successive steps of the E-M algorithm can be described as follows:

Starting with initial values of the of haplotypes frequencies, noted  $p_1^{(0)}, p_2^{(0)}, \dots, p_H^{(0)}$ , these initial values serve to estimate the genotype frequencies (Expectation step) and then estimate another set of haplotype frequencies (Maximisation step), noted  $p_1^{(1)}, p_2^{(1)}, \dots, p_H^{(1)}$ ,  
 30 these two steps are iterated until changes in the sets of haplotypes frequency are very small.

A stop criterion can be that the maximum difference between haplotype frequencies between two iterations is less than  $10^{-7}$ . These values can be adjusted according to the desired precision of estimations.

In details, at a given iteration  $s$ , the Expectation step consists in calculating the  
5 genotypes frequencies by the following equation:

$$\begin{aligned} pr(genotype_i)^{(s)} &= pr(phenotype_j) \cdot pr(genotype_i | phenotype_j)^{(s)} \\ &= \frac{n_j}{N} \cdot \frac{pr(h_k, h_l)^{(s)}}{p_j^{(s)}} \end{aligned} \quad \text{Equation 3}$$

where genotype  $i$  occurs in phenotype  $j$ , and where  $h_k$  and  $h_l$  constitute genotype  $i$ . Each probability is derived according to eq.1, and eq.2 described above.

Then the Maximisation step simply estimates another set of haplotype frequencies  
10 given the genotypes frequencies. This approach is also known as gene-counting method (Smith, *Ann. Hum. Genet.*, 21:254-276, 1957).

$$p_i^{(s+1)} = \frac{1}{2} \sum_{j=1}^F \sum_{l=1}^{c_j} \delta_{il} \cdot pr(genotype_i)^{(s)} \quad \text{Equation 4}$$

Where  $\delta_{il}$  is an indicator variable which count the number of time haplotype  $l$  in genotype  $i$ . It takes the values of 0, 1 or 2.

15 To ensure that the estimation finally obtained is the maximum-likelihood estimation several values of departures are required. The estimations obtained are compared and if they are different the estimations leading to the best likelihood are kept.

### 3) Methods to calculate linkage disequilibrium between markers

A number of methods can be used to calculate linkage disequilibrium between any  
20 two genetic positions, in practice linkage disequilibrium is measured by applying a statistical association test to haplotype data taken from a population.

Linkage disequilibrium between any pair of biallelic markers comprising at least one of the biallelic markers of the present invention ( $M_i, M_j$ ) having alleles ( $a_i/b_i$ ) at marker  $M_i$  and alleles ( $a_j/b_j$ ) at marker  $M_j$  can be calculated for every allele combination ( $a_i, a_j, a_i, b_j, b_i, a_j$   
25 and  $b_i, b_j$ ), according to the Piazza formula :

$$\Delta_{aiaj} = \sqrt{\theta 4} - \sqrt{(\theta 4 + \theta 3)(\theta 4 + \theta 2)}, \text{ where :}$$

$\theta 4 = - -$  = frequency of genotypes not having allele  $a_i$  at  $M_i$  and not having allele  $a_j$  at  $M_j$

$\theta 3 = - +$  = frequency of genotypes not having allele  $a_i$  at  $M_i$  and having allele  $a_j$  at  $M_j$

30  $\theta 2 = + -$  = frequency of genotypes having allele  $a_i$  at  $M_i$  and not having allele  $a_j$  at  $M_j$

Linkage disequilibrium (LD) between pairs of biallelic markers ( $M_i, M_j$ ) can also be calculated for every allele combination ( $a_i, a_j; a_i, b_j; b_i, a_j$  and  $b_i, b_j$ ), according to the maximum-likelihood estimate (MLE) for delta (the composite genotypic disequilibrium coefficient), as described by Weir (Weir B.S., *Genetic Data Analysis, Sinauer Ass. Eds,*

5 1996). The MLE for the composite linkage disequilibrium is:

$$D_{aiaj} = (2n_1 + n_2 + n_3 + n_4/2)/N - 2(\text{pr}(a_i) \cdot \text{pr}(a_j))$$

Where  $n_1 = \Sigma$  phenotype ( $a_i/a_i, a_j/a_j$ ),  $n_2 = \Sigma$  phenotype ( $a_i/a_i, a_j/b_j$ ),  $n_3 = \Sigma$  phenotype ( $a_i/b_i, a_j/a_j$ ),  $n_4 = \Sigma$  phenotype ( $a_i/b_i, a_j/b_j$ ) and  $N$  is the number of individuals in the sample.

This formula allows linkage disequilibrium between alleles to be estimated when  
10 only genotype, and not haplotype, data are available.

Another means of calculating the linkage disequilibrium between markers is as follows. For a couple of biallelic markers,  $M_i (a/b_i)$  and  $M_j (a/b_j)$ , fitting the Hardy-Weinberg equilibrium, one can estimate the four possible haplotype frequencies in a given population according to the approach described above.

15 The estimation of gametic disequilibrium between  $a_i$  and  $a_j$  is simply:

$$D_{aiaj} = \text{pr}(\text{haplotype}(a_i, a_j)) - \text{pr}(a_i) \cdot \text{pr}(a_j).$$

Where  $\text{pr}(a_i)$  is the probability of allele  $a_i$  and  $\text{pr}(a_j)$  is the probability of allele  $a_j$  and where  $\text{pr}(\text{haplotype}(a_i, a_j))$  is estimated as in Equation 3 above.

For a couple of biallelic marker only one measure of disequilibrium is necessary to  
20 describe the association between  $M_i$  and  $M_j$ .

Then a normalised value of the above is calculated as follows:

$$D'_{aiaj} = D_{aiaj} / \max (-\text{pr}(a_i) \cdot \text{pr}(a_j), -\text{pr}(b_i) \cdot \text{pr}(b_j)) \text{ with } D_{aiaj} < 0$$

$$D'_{aiaj} = D_{aiaj} / \max (\text{pr}(b_i) \cdot \text{pr}(a_j), \text{pr}(a_i) \cdot \text{pr}(b_j)) \text{ with } D_{aiaj} > 0$$

The skilled person will readily appreciate that other LD calculation methods can be  
25 used without undue experimentation.

Linkage disequilibrium among a set of biallelic markers having an adequate heterozygosity rate can be determined by genotyping between 50 and 1000 unrelated individuals, preferably between 75 and 200, more preferably around 100.

#### 4) Testing for association

30 Methods for determining the statistical significance of a correlation between a phenotype and a genotype, in this case an allele at a biallelic marker or a haplotype made up of such alleles, may be determined by any statistical test known in the art and with any accepted threshold of statistical significance being required. The application of particular

methods and thresholds of significance are well within the skill of the ordinary practitioner of the art.

Testing for association is performed by determining the frequency of a biallelic marker allele in case and control populations and comparing these frequencies with a statistical test to determine if there is a statistically significant difference in frequency which would indicate a correlation between the trait and the biallelic marker allele under study. Similarly, a haplotype analysis is performed by estimating the frequencies of all possible haplotypes for a given set of biallelic markers in case and control populations, and comparing these frequencies with a statistical test to determine if there is a statistically significant correlation between the haplotype and the phenotype (trait) under study. Any statistical tool useful to test for a statistically significant association between a genotype and a phenotype may be used. Preferably the statistical test employed is a chi-square test with one degree of freedom. A P-value is calculated (the P-value is the probability that a statistic as large or larger than the observed one would occur by chance).

#### 15 **Statistical significance**

In preferred embodiments, significance for diagnosis purposes, either as a positive basis for further diagnostic tests or as a preliminary starting point for early preventive therapy, the p value related to a biallelic marker association is preferably about  $1 \times 10^{-2}$  or less, more preferably about  $1 \times 10^{-4}$  or less, for a single biallelic marker analysis and about  $1 \times 10^{-3}$  or less, still more preferably  $1 \times 10^{-6}$  or less and most preferably of about  $1 \times 10^{-8}$  or less, for a haplotype analysis involving several markers. These values are believed to be applicable to any association studies involving single or multiple marker combinations.

The skilled person can use the range of values set forth above as a starting point in order to carry out association studies with biallelic markers of the present invention. In doing so, significant associations between the biallelic markers of the present invention and responses to drugs or side effects upon treatment with drugs or diseases involving the metabolic conversion of xenobiotics can be revealed and used for diagnosis and drug screening purposes.

#### **Phenotypic permutation**

30 In order to confirm the statistical significance of the first stage haplotype analysis described above, it might be suitable to perform further analyses in which genotyping data from case-control individuals are pooled and randomised with respect to the trait phenotype. Each individual genotyping data is randomly allocated to two groups, which contain the same number of individuals as the case-control populations used to compile the data

obtained in the first stage. A second stage haplotype analysis is preferably run on these artificial groups, preferably for the markers included in the haplotype of the first stage analysis showing the highest relative risk coefficient. This experiment is reiterated preferably at least between 100 and 10000 times. The repeated iterations allow the

5 determination of the percentage of obtained haplotypes with a significant p-value level.

#### Assessment of statistical association

To address the problem of false positives similar analysis may be performed with the same case-control populations in random genomic regions. Results in random regions and the candidate region are compared as described in US Provisional Patent Application

10 entitled "Methods, software and apparatus for identifying genomic regions harbouring a gene associated with a detectable trait".

#### 5) Evaluation of risk factors

The association between a risk factor (in genetic epidemiology the risk factor is the presence or the absence of a certain allele or haplotype at marker loci) and a disease is

15 measured by the odds ratio (OR) and by the relative risk (RR). If  $P(R^+)$  is the probability of developing the disease for individuals with R and  $P(R^-)$  is the probability for individuals without the risk factor, then the relative risk is simply the ratio of the two probabilities, that is:

$$RR = P(R^+)/P(R^-)$$

20 In case-control studies, direct measures of the relative risk cannot be obtained because of the sampling design. However, the odds ratio allows a good approximation of the relative risk for low-incidence diseases and can be calculated:

$$OR = \left[ \frac{F^+}{1 - F^+} \right] / \left[ \frac{F^-}{(1 - F^-)} \right]$$

$F^+$  is the frequency of the exposure to the risk factor in cases and  $F^-$  is the frequency

25 of the exposure to the risk factor in controls.  $F^+$  and  $F^-$  are calculated using the allelic or haplotype frequencies of the study and further depend on the underlying genetic model (dominant, recessive, additive...).

One can further estimate the attributable risk (AR) which describes the proportion of individuals in a population exhibiting a trait due to a given risk factor. This measure is

30 important in quantitating the role of a specific factor in disease etiology and in terms of the public health impact of a risk factor. The public health relevance of this measure lies in

estimating the proportion of cases of disease in the population that could be prevented if the exposure of interest were absent. AR is determined as follows:

$$AR = P_E (RR-1) / (P_E (RR-1)+1)$$

AR is the risk attributable to a biallelic marker allele or a biallelic marker haplotype.

- 5  $P_E$  is the frequency of exposure to an allele or a haplotype within the population at large; and RR is the relative risk which, is approximated with the odds ratio when the trait under study has a relatively low incidence in the general population.

#### **IV.D. Identification of Biallelic Markers in Linkage Disequilibrium with the Biallelic Markers of the Invention**

- 10 Once a first biallelic marker has been identified in a genomic region of interest, the practitioner of ordinary skill in the art, using the teachings of the present invention, can easily identify additional biallelic markers in linkage disequilibrium with this first marker. As mentioned before any marker in linkage disequilibrium with a first marker associated with a trait will be associated with the trait. Therefore, once an association has been  
15 demonstrated between a given biallelic marker and a trait, the discovery of additional biallelic markers associated with this trait is of great interest in order to increase the density of biallelic markers in this particular region. The causal gene or mutation will be found in the vicinity of the marker or set of markers showing the highest correlation with the trait.

- Identification of additional markers in linkage disequilibrium with a given marker  
20 involves: (a) amplifying a genomic fragment comprising a first biallelic marker from a plurality of individuals; (b) identifying of second biallelic markers in the genomic region harboring said first biallelic marker; (c) conducting a linkage disequilibrium analysis between said first biallelic marker and second biallelic markers; and (d) selecting said second biallelic markers as being in linkage disequilibrium with said first marker.

- 25 Subcombinations comprising steps (b) and (c) are also contemplated.

- Methods to identify biallelic markers and to conduct linkage disequilibrium analysis are described herein and can be carried out by the skilled person without undue experimentation. The present invention then also concerns biallelic markers which are in linkage disequilibrium with the specific biallelic markers shown in Figure 1 and which are  
30 expected to present similar characteristics in terms of their respective association with a given trait.

#### **IV.E. Identification of Functional Mutations**

Once a positive association is confirmed with a biallelic marker of the present invention, the associated candidate gene can be scanned for mutations by comparing the



sequences of a selected number of trait positive and trait negative or control individuals. In a preferred embodiment, functional regions such as exons and splice sites, promoters and other regulatory regions of the candidate gene are scanned for mutations. Preferably, trait positive individuals carry the haplotype shown to be associated with the trait and trait negative individuals do not carry the haplotype or allele associated with the trait. The mutation detection procedure is essentially similar to that used for biallelic site identification.

The method used to detect such mutations generally comprises the following steps: (a) amplification of a region of the candidate gene comprising a biallelic marker or a group of biallelic markers associated with the trait from DNA samples of trait positive patients and trait negative controls; (b) sequencing of the amplified region; (c) comparison of DNA sequences from trait-positive patients and trait-negative controls; and (d) determination of mutations specific to trait-positive patients. Subcombinations which comprise steps (b) and (c) are specifically contemplated.

It is preferred that candidate polymorphisms be then verified by screening a larger population of cases and controls by means of any genotyping procedure such as those described herein, preferably using a microsequencing technique in an individual test format. Polymorphisms are considered as candidate mutations when present in cases and controls at frequencies compatible with the expected association results.

Identification of mutations and low frequency polymorphisms in exons 3-5, in the 5'UTR region and in the 3' flanking region of the MGST-II gene is further described in Example 5. Eight polymorphisms were identified in the region of the MGST-II gene that was scanned. Three mutations were identified in the 3'UTR region. One mutation in exon 4 causes an amino acid substitution (Tyr→His) at the polypeptide level. A mutation in exon 5 introduces a stop codon into the ORF leading to the expression of a truncated MGST-II polypeptide. These mutations modify the specificity, activity and function of the MGST-II enzyme and therefore represent functional mutations of the MGST-II gene. Candidate polymorphisms and mutations suspected of being responsible for the detectable phenotype, such as hepatotoxicity to zileuton or asthma, can be confirmed by screening a larger population of affected and unaffected individuals using any of the genotyping procedures described herein. Preferably the microsequencing technique is used. In a preferred embodiment trait positive and trait negative populations are genotyped for the candidate polymorphisms identified in Example 5 (10-286-289, 10-286-345, 10-286-375, 10-523-232, 10-289-201, 10-290-37, 10-290-326 and 10-290-328) most preferably for the

mutations in exons 4 and 5 (10-289-201 and 10-290-37). Polymorphisms are considered as candidate "trait-causing" mutations when they exhibit a statistically significant correlation with the detectable phenotype.

**V. Biallelic Markers of the Invention in Methods of Genetic Diagnostics**

5        The biallelic markers of the present invention can also be used to develop diagnostics tests capable of identifying individuals who express a detectable trait as the result of a specific genotype or individuals whose genotype places them at risk of developing a detectable trait at a subsequent time. The trait analyzed using the present diagnostics may be any detectable trait, including a response to a drug or side effects to a  
10 drug upon treatment or a disease involving the metabolic conversion of xenobiotics.

      The diagnostic techniques of the present invention may employ a variety of methodologies to determine whether a test subject has a biallelic marker pattern associated with an increased risk of developing a detectable trait or whether the individual suffers from a detectable trait as a result of a particular mutation, including methods which enable the  
15 analysis of individual chromosomes for haplotyping, such as family studies, single sperm DNA analysis or somatic hybrids.

      The present invention provides diagnostic methods to determine whether an individual is at risk of developing a disease or suffers from a disease resulting from a mutation or a polymorphism in a candidate gene of the present invention. The present  
20 invention also provides methods to determine whether an individual is likely to respond positively to a therapeutic agent or whether an individual is at risk of developing an adverse side effect to a therapeutic agent.

      These methods involve obtaining a nucleic acid sample from the individual and, determining, whether the nucleic acid sample contains at least one allele or at least one  
25 biallelic marker haplotype, indicative of a risk of developing the trait or indicative that the individual expresses the trait as a result of possessing a particular candidate gene polymorphism or mutation (trait-causing allele).

      Preferably, in such diagnostic methods, a nucleic acid sample is obtained from the individual and this sample is genotyped using methods described above in III. "Methods of  
30 Genotyping an Individual for Biallelic Markers." The diagnostics may be based on a single biallelic marker or a on group of biallelic markers.

      In each of these methods, a nucleic acid sample is obtained from the test subject and the biallelic marker pattern of one or more of the biallelic markers listed in Figure 1 is determined.

In one embodiment, a PCR amplification is conducted on the nucleic acid sample to amplify regions in which polymorphisms associated with a detectable phenotype have been identified. The amplification products are sequenced to determine whether the individual possesses one or more polymorphisms associated with a detectable phenotype. The primers  
5 used to generate amplification products may comprise the primers listed in Figure 7. Alternatively, the nucleic acid sample is subjected to microsequencing reactions as described above to determine whether the individual possesses one or more polymorphisms associated with a detectable phenotype resulting from a mutation or a polymorphism in a candidate gene. The primers used in the microsequencing reactions may include the primers  
10 listed in Figure 6. In another embodiment, the nucleic acid sample is contacted with one or more allele specific oligonucleotide probes which, specifically hybridize to one or more candidate gene alleles associated with a detectable phenotype. The probes used in the hybridization assay may include the probes listed in Figure 8. Diagnostic kits comprising polynucleotides of the present invention are further described in section I.

15 These diagnostic methods are extremely valuable as they can, in certain circumstances, be used to initiate preventive treatments or to allow an individual carrying a significant haplotype to foresee warning signs such as minor symptoms. For diseases in which attacks may be extremely violent and sometimes fatal if not treated on time, the knowledge of a potential predisposition, even if this predisposition is not absolute, might  
20 contribute in a very significant manner to treatment efficacy. Similarly, a diagnosed predisposition to a potential side effect could immediately direct the physician toward a treatment for which such side effects have not been observed during clinical trials.

Diagnostics, which analyze and predict response to a drug or side effects to a drug, may be used to determine whether an individual should be treated with a particular drug.  
25 For example, if the diagnostic indicates a likelihood that an individual will respond positively to treatment with a particular drug, the drug may be administered to the individual. Conversely, if the diagnostic indicates that an individual is likely to respond negatively to treatment with a particular drug, an alternative course of treatment may be prescribed. A negative response may be defined as either the absence of an efficacious  
30 response or the presence of toxic side effects.

Clinical drug trials represent another application for the markers of the present invention. One or more markers indicative of response to a drug or to side effects to a drug may be identified using the methods described above. Thereafter, potential participants in clinical trials of such an agent may be screened to identify those individuals most likely to

respond favorably to the drug and exclude those likely to experience side effects. In that way, the effectiveness of drug treatment may be measured in individuals who respond positively to the drug, without lowering the measurement as a result of the inclusion of individuals who are unlikely to respond positively in the study and without risking  
 5 undesirable safety problems.

In a preferred embodiment the identity of the nucleotide present at, at least one, biallelic marker selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-  
 10 219, 12-721-440 and 10-420-284 is determined, and optionally wherein the detectable trait is asthma, or optionally the detectable trait is hepatotoxicity to the anti-asthmatic drug zileuton. In another preferred embodiment the identity of the nucleotide present at, at least one of the polymorphic sites selected from the group consisting of 12-447-58, 12-455-326, 12-461-299, 12-453-429, 12-424-198, 12-454-363, 12-716-295, 10-428-219, 12-720-80, 10-  
 15 420-248, 12-721-440, 12-653-423, 10-470-25, 10-471-84, 10-471-85, 12-637-219 and 12-652-203 is determined, and optionally wherein the detectable trait is asthma. In another preferred embodiment the identity of the nucleotide present at, at least one of the polymorphic sites selected from the group consisting of 12-453-429, 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-  
 20 219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284 is determined, and optionally wherein the detectable trait is hepatotoxicity to the anti-asthmatic drug zileuton. Diagnostic kits comprising polynucleotides of the present invention are further described in section I.  
 "Biallelic Markers and Polynucleotides Comprising Biallelic Markers."

## 25 **VI. Association of Biallelic Markers of the Invention With Asthma**

In the context of the present invention, an association between the MGST-II, ME1, UGT1A7 and UGT2B4 genes and asthma was established.

Asthma affects over 5% of the population in industrialized countries. It is increasing in prevalence and severity and has a rising mortality (Rang H.P., Ritter J.M. and  
 30 Dale M.M.; *Pharmacology*; Churchill Livingstone, NY, 1995). Bronchial asthma is a multifactorial syndrome rather than a single disease, defined as airway obstruction characterized by inflammatory changes in the airways and bronchial hyper-responsiveness. In addition to the evidenced impact of environmental factors on the development of asthma, patterns of clustering and segregation in asthmatic families have suggested a genetic

component to asthma. However the lack of a defined and specific asthma phenotype and of suitable markers for genetic analysis is proving to be a major hurdle for reliably identifying genes associated with asthma. The identification of genes implicated in asthma would represent a major step towards the identification of new molecular targets for the development of anti-asthma drugs. Moreover there is no straightforward physiological or biological blood test for the asthmatic state. As a result, adequate asthma treatment is often delayed, thereby allowing the inflammation process to better establish itself. Thus, there is a need for the identification of asthma susceptibility genes in order to develop an efficient and reliable asthma diagnostic test.

As mentioned above, products of arachidonic acid metabolism are important inflammatory mediators and have been involved in a number of inflammatory diseases, including asthma. More specifically, prostaglandins and leukotrienes are thought to play a major role in the inflammatory process observed in asthma patients.

In order to investigate and identify a genetic origin to asthma, a candidate gene scan was conducted. This approach comprised:

- selecting candidate genes potentially involved in the pathological pathway of interest, in this case arachidonic acid metabolism, and
- identifying biallelic markers in those genes, and finally
- conducting association studies to identify biallelic marker alleles or haplotypes associated with asthma.

Further details concerning this association study are provided in Example 3, results are briefly summarized below.

Two groups of independent individuals were used in this association study in accordance with the invention: the case-control populations. The two groups corresponded to 297 asthmatic individuals and 178 control individuals. The trait positive asthma population was mostly composed of individuals from Caucasian ethnic background (>90 %). The control population was composed of individuals from a random US Caucasian population.

In the association study described in Example 3, several biallelic marker haplotypes were shown to be significantly associated with asthma. A preferred haplotype consisting of three biallelic markers (12-455-326, 12-453-429 and 12-424-198 ) presented a p-value of  $3.2 \cdot 10^{-5}$ . Another preferred haplotype consisting of four biallelic markers (12-455-326, 12-453-429, 12-424-198 and 12-454-363) had a p-value of  $1.2 \cdot 10^{-6}$ . Phenotypic permutation

tests confirmed the statistical significance of these results. These haplotypes can therefore be considered to be significantly associated with asthma.

This information is extremely valuable. The knowledge of a potential genetic predisposition to asthma, even if this predisposition is not absolute, might contribute in a very significant manner to treatment efficacy of asthma patients and to the development of new therapeutic and diagnostic tools.

**VII. Association of Biallelic Markers of the Invention with Hepatotoxicity to Anti-Asthma Drug Zileuton (Zyflo™)**

In the context of the present invention, an association between the MGST-II gene and side effects related to treatment with the anti-asthmatic drug zileuton was discovered.

As mentioned above, bronchial asthma is a multifactorial syndrome rather than a single disease, defined as airway obstruction characterized by inflammatory changes in the airways and bronchial hyper-responsiveness. Although initially reversible with bronchodilators, airway obstruction becomes increasingly irreversible if treated poorly. Asthma management therefore relies on early and regular use of drugs that control the disease. As a consequence, there is a strong need for efficient and safe therapeutic opportunities for patients with asthma. There are two main categories of anti-asthmatic drugs— bronchodilators and anti-inflammatory agents. There is now general agreement on the need to implement early anti-inflammatory treatment rather than relying on symptomatic treatment with bronchodilators alone. The leukotrienes, a family of proinflammatory mediators arising via arachidonic acid metabolism, have been implicated in the inflammatory cascade that occurs in asthmatic airways. Of great relevance to the pathogenesis of asthma are the 5-lipoxygenase and the 5-lipoxygenase activating protein (FLAP), which catalyze the initial steps in the biosynthesis of leukotrienes from arachidonic acid. Given the significant role of the inflammatory process in asthma, pharmacological agents, such as leukotriene antagonists, FLAP inhibitors and 5-lipoxygenase inhibitors have been developed.

Zileuton (Zyflo™) is an active inhibitor of 5-lipoxygenase, the enzyme that catalyzes the formation of leukotrienes from arachidonic acid, indicated for prophylaxis and chronic treatment of asthma. A minority of zileuton-treated patients develop liver function abnormalities as close monitoring revealed that elevations of liver function tests may occur during treatment with zileuton. In the present invention the ALT test (serum level of alanine aminotransferase) was used, which is considered the most sensitive indicator of liver injury.

In order to investigate and identify a genetic origin to zileuton-associated hepatotoxicity, a candidate gene scan was conducted. This approach comprised:

- selecting candidate genes potentially involved in the pathological pathway of interest or in the metabolism of zileuton, and
- 5 - identifying biallelic markers in those genes, and finally
- conducting association studies to identify biallelic marker alleles or haplotypes associated with elevations of liver function tests upon treatment with zileuton.

Further details concerning this association study are provided in Example 4, results are briefly summarized below.

- 10 Two groups of unrelated individuals were used in this association study in accordance with the invention: the case-control populations. The case population was composed of 89 asthmatic individuals treated with zileuton showing zileuton-associated hepatotoxicity monitored by the serum level of alanine aminotransferase (ALT) and the control population was composed of 208 asthmatic individuals treated with zileuton and
- 15 having no increased serum ALT level.

The association study conducted with the biallelic markers derived from the MGST-II locus showed that several haplotypes were significantly associated with zileuton-associated hepatotoxicity. A preferred haplotype consisting of three biallelic markers (12-441-233, 12-461-299 and 12-453-429) presented a p-value of  $1.5 \cdot 10^{-5}$  and an odd ratio of

20 3.63. A second preferred haplotype consisting of four biallelic markers (12-441-233, 12-461-299, 12-453-429 and 12-426-154) had a p-value of  $5.2 \cdot 10^{-7}$  and an odd ratio of 5.75.

This information is extremely valuable. The knowledge of a potential genetic predisposition to hepatotoxicity upon treatment with zileuton, even if this predisposition is not absolute, might contribute in a very significant manner to the safety of asthma treatment and

25 to the development of diagnostic tools.

Similar association studies, with different case-control populations, can be routinely carried out by the skilled technician using the biallelic markers of the present invention in order to identify other association between traits and MGST-II-related biallelic marker alleles or haplotypes.

## 30 **VI. Computer-Related Embodiments**

As used herein the term "nucleic acid codes of the invention" encompass the nucleotide sequences comprising, consisting essentially of, or consisting of any one of the following: a) a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500 or 1000 nucleotides, to the extent that a polynucleotide of these lengths

is consistent with the lengths of the particular Sequence ID, of a sequence selected from the group consisting of the sequences described in Figure 2, and the complements thereof; b) a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 500 or 1000 nucleotides, to the extent that a polynucleotide of these lengths is consistent

5 with the lengths of the particular Sequence ID, of a sequence selected from the group consisting of the sequences described in Figure 3, and the complements thereof; c) a contiguous span of at least 12, 15, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, or 500 nucleotides, to the extent that a polynucleotide of these lengths is consistent with the lengths of the particular Sequence ID, of a sequence selected from the group consisting of

10 the sequences described in Figure 6, more preferably a set of markers or sequences consisting of those markers or sequences found in SEQ ID Nos. 3, 5, 9, 13-15, 25, 31, 33, 37, 38, 41, 323, 345, 351-353, 357, 377, and 380, and the complements thereof, wherein said span includes an DME-related biallelic marker, preferably an DME-related biallelic marker described in Figure 1, preferably the biallelic markers found in Figures 9, 10, 11 and

15 12; or more preferably the biallelic markers found in SEQ ID Nos. 3, 5, 9, 13-15, 25, 31, 33, 37, 38, 41, 323, 345, 351-353, 357, 377, 380, in said sequence with the alternative allele present at said biallelic marker.

The "nucleic acid codes of the invention" further encompass nucleotide sequences homologous to a contiguous span of at least 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200,

20 500 or 1000 nucleotides, to the extent that a contiguous span of these lengths is consistent with the lengths of the particular Sequence ID, of a sequence selected from the group consisting of the sequences described in Figure 2, Figure 3 and Figure 6 and the complements thereof. Homologous sequences refer to a sequence having at least 99%, 98%, 97%, 96%, 95%, 90%, 85%, 80%, or 75% homology to these contiguous spans. Homology

25 may be determined using any method described herein, including BLAST2N with the default parameters or with any modified parameters. Homologous sequences also may include RNA sequences in which uridines replace the thymines in the nucleic acid codes of the invention. It will be appreciated that the nucleic acid codes of the invention can be represented in the traditional single character format (See the inside back cover of Stryer, Lubert. *Biochemistry*,

30 3<sup>rd</sup> edition. W. H Freeman & Co., New York.) or in any other format or code which records the identity of the nucleotides in a sequence.

It will be appreciated by those skilled in the art that the nucleic acid codes of the invention, one or more of the polypeptide codes of SEQ ID Nos. 488 and 489 can be stored, recorded, and manipulated on any medium which can be read and accessed by a computer. As



used herein, the words "recorded" and "stored" refer to a process for storing information on a computer medium. A skilled artisan can readily adopt any of the presently known methods for recording information on a computer readable medium to generate manufactures comprising one or more of the nucleic acid codes of the invention and one or more of the polypeptide

5 codes of SEQ ID Nos. 488 and 489. Another aspect of the present invention is a computer readable medium having recorded thereon at least 2, 5, 10, 15, 20, 25, 30, or 50 nucleic acid codes of the invention, and the complements thereof. Another aspect of the present invention is a computer readable medium having recorded thereon at least 2, 5, 10, 15, 20, 25, 30, or 50 polypeptide codes of SEQ ID Nos. 488 and 489.

10 Computer readable media include magnetically readable media, optically readable media, electronically readable media and magnetic/optical media. For example, the computer readable media may be a hard disk, a floppy disk, a magnetic tape, CD-ROM, Digital Versatile Disk (DVD), Random Access Memory (RAM), or Read Only Memory (ROM) as well as other types of other media known to those skilled in the art.

15 Embodiments of the present invention include systems, particularly computer systems which store and manipulate the sequence information described herein. One example of a computer system 100 is illustrated in block diagram form in Figure 17. As used herein, "a computer system" refers to the hardware components, software components, and data storage components used to analyze the nucleotide sequences of the nucleic acid codes of the invention

20 , or the amino acid sequences of the polypeptide codes of SEQ ID Nos. 488 and 489. In one embodiment, the computer system 100 is a Sun Enterprise 1000 server (Sun Microsystems, Palo Alto, CA). The computer system 100 preferably includes a processor for processing, accessing and manipulating the sequence data. The processor 105 can be any well-known type of central processing unit, such as the Pentium III from Intel Corporation, or similar processor

25 from Sun, Motorola, Compaq or International Business Machines. Preferably, the computer system 100 is a general purpose system that comprises the processor 105 and one or more internal data storage components 110 for storing data, and one or more data retrieving devices for retrieving the data stored on the data storage components. A skilled artisan can readily appreciate that any one of the currently available computer systems are suitable. In one

30 particular embodiment, the computer system 100 includes a processor 105 connected to a bus which is connected to a main memory 115 (preferably implemented as RAM) and one or more internal data storage devices 110, such as a hard drive and/or other computer readable media having data recorded thereon. In some embodiments, the computer system 100 further includes one or more data retrieving device 118 for reading the data stored on the internal data

storage devices 110. The data retrieving device 118 may represent, for example, a floppy disk drive, a compact disk drive, a magnetic tape drive, etc. In some embodiments, the internal data storage device 110 is a removable computer readable medium such as a floppy disk, a compact disk, a magnetic tape, etc. containing control logic and/or data recorded thereon. The computer system 100 may advantageously include or be programmed by appropriate software for reading the control logic and/or the data from the data storage component once inserted in the data retrieving device. The computer system 100 includes a display 120 which is used to display output to a computer user. It should also be noted that the computer system 100 can be linked to other computer systems 125a-c in a network or wide area network to provide centralized access to the computer system 100. Software for accessing and processing the nucleotide sequences of the nucleic acid codes of the invention, or the amino acid sequences of the polypeptide codes of SEQ ID Nos. 488 and 489 (such as search tools, compare tools, and modeling tools etc.) may reside in main memory 115 during execution. In some embodiments, the computer system 100 may further comprise a sequence comparer for comparing the above-described nucleic acid codes of the invention or polypeptide codes of SEQ ID Nos. 488 and 489 stored on a computer readable medium to reference nucleotide or polypeptide sequences stored on a computer readable medium. A "sequence comparer" refers to one or more programs which are implemented on the computer system 100 to compare a nucleotide or polypeptide sequence with other nucleotide or polypeptide sequences and/or compounds including but not limited to peptides, peptidomimetics, and chemicals stored within the data storage means. For example, the sequence comparer may compare the nucleotide sequences of the nucleic acid codes of the invention, or the amino acid sequences of the polypeptide codes of SEQ ID Nos. 488 and 489 stored on a computer readable medium to reference sequences stored on a computer readable medium to identify homologies, motifs implicated in biological function, or structural motifs. The various sequence comparer programs identified elsewhere in this patent specification are particularly contemplated for use in this aspect of the invention.

Figure 18 is a flow diagram illustrating one embodiment of a process 200 for comparing a new nucleotide or protein sequence with a database of sequences in order to determine the homology levels between the new sequence and the sequences in the database. The database of sequences can be a private database stored within the computer system 100, or a public database such as GENBANK, PIR OR SWISSPROT that is available through the Internet.

The process 200 begins at a start state 201 and then moves to a state 202 wherein the new sequence to be compared is stored to a memory in a computer system 100. As discussed

above, the memory could be any type of memory, including RAM or an internal storage device. The process 200 then moves to a state 204 wherein a database of sequences is opened for analysis and comparison. The process 200 then moves to a state 206 wherein the first sequence stored in the database is read into a memory on the computer. A comparison is then

5 performed at a state 210 to determine if the first sequence is the same as the second sequence. It is important to note that this step is not limited to performing an exact comparison between the new sequence and the first sequence in the database. Well-known methods are known to those of skill in the art for comparing two nucleotide or protein sequences, even if they are not identical. For example, gaps can be introduced into one sequence in order to raise the

10 homology level between the two tested sequences. The parameters that control whether gaps or other features are introduced into a sequence during comparison are normally entered by the user of the computer system. Once a comparison of the two sequences has been performed at the state 210, a determination is made at a decision state 210 whether the two sequences are the same. Of course, the term "same" is not limited to sequences that are absolutely identical.

15 Sequences that are within the homology parameters entered by the user will be marked as "same" in the process 200. If a determination is made that the two sequences are the same, the process 200 moves to a state 214 wherein the name of the sequence from the database is displayed to the user. This state notifies the user that the sequence with the displayed name fulfills the homology constraints that were entered. Once the name of the stored sequence is

20 displayed to the user, the process 200 moves to a decision state 218 wherein a determination is made whether more sequences exist in the database. If no more sequences exist in the database, then the process 200 terminates at an end state 220. However, if more sequences do exist in the database, then the process 200 moves to a state 224 wherein a pointer is moved to the next sequence in the database so that it can be compared to the new sequence. In this

25 manner, the new sequence is aligned and compared with every sequence in the database. It should be noted that if a determination had been made at the decision state 212 that the sequences were not homologous, then the process 200 would move immediately to the decision state 218 in order to determine if any other sequences were available in the database for comparison. Accordingly, one aspect of the present invention is a computer system

30 comprising a processor, a data storage device having stored thereon a nucleic acid code of the invention or a polypeptide code of SEQ ID Nos. 488 and 489, a data storage device having retrievably stored thereon reference nucleotide sequences or polypeptide sequences to be compared to the nucleic acid code of the invention or polypeptide code of SEQ ID Nos. 488 and 489 and a sequence comparer for conducting the comparison. The sequence

comparer may indicate a homology level between the sequences compared or identify structural motifs in the above described nucleic acid code of the invention and polypeptide codes of SEQ ID Nos. 488 and 489 or it may identify structural motifs in sequences which are compared to these nucleic acid codes and polypeptide codes. In some embodiments, the data storage device may have stored thereon the sequences of at least 2, 5, 10, 15, 20, 25, 30, or 50 of the nucleic acid codes of the invention or polypeptide codes of SEQ ID Nos. 488 and 489.

Another aspect of the present invention is a method for determining the level of homology between a nucleic acid code of the invention and a reference nucleotide sequence, comprising the steps of reading the nucleic acid code and the reference nucleotide sequence through the use of a computer program which determines homology levels and determining homology between the nucleic acid code and the reference nucleotide sequence with the computer program. The computer program may be any of a number of computer programs for determining homology levels, including those specifically enumerated herein, including BLAST2N with the default parameters or with any modified parameters. The method may be implemented using the computer systems described above. The method may also be performed by reading 2, 5, 10, 15, 20, 25, 30, or 50 of the above described nucleic acid codes of the invention through use of the computer program and determining homology between the nucleic acid codes and reference nucleotide sequences .

Figure 19 is a flow diagram illustrating one embodiment of a process 250 in a computer for determining whether two sequences are homologous. The process 250 begins at a start state 252 and then moves to a state 254 wherein a first sequence to be compared is stored to a memory. The second sequence to be compared is then stored to a memory at a state 256. The process 250 then moves to a state 260 wherein the first character in the first sequence is read and then to a state 262 wherein the first character of the second sequence is read. It should be understood that if the sequence is a nucleotide sequence, then the character would normally be either A, T, C, G or U. If the sequence is a protein sequence, then it should be in the single letter amino acid code so that the first and sequence sequences can be easily compared.

A determination is then made at a decision state 264 whether the two characters are the same. If they are the same, then the process 250 moves to a state 268 wherein the next characters in the first and second sequences are read. A determination is then made whether the next characters are the same. If they are, then the process 250 continues this loop until two characters are not the same. If a determination is made that the next two

characters are not the same, the process 250 moves to a decision state 274 to determine whether there are any more characters either sequence to read. If there aren't any more characters to read, then the process 250 moves to a state 276 wherein the level of homology between the first and second sequences is displayed to the user. The level of homology is  
5 determined by calculating the proportion of characters between the sequences that were the same out of the total number of sequences in the first sequence. Thus, if every character in a first 100 nucleotide sequence aligned with a every character in a second sequence, the homology level would be 100%. Alternatively, the computer program may be a computer program which compares the nucleotide sequences of the nucleic acid codes of the present  
10 invention, to reference nucleotide sequences in order to determine whether the nucleic acid code of SEQ ID Nos. 1-652 differs from a reference nucleic acid sequence at one or more positions. Optionally such a program records the length and identity of inserted, deleted or substituted nucleotides with respect to the sequence of either the reference polynucleotide or the nucleic acid code of SEQ ID Nos. 1-652. In one embodiment, the computer program may  
15 be a program which determines whether the nucleotide sequences of the nucleic acid codes of the invention contain a biallelic marker or single nucleotide polymorphism (SNP) with respect to a reference nucleotide sequence. This single nucleotide polymorphism may comprise a single base substitution, insertion, or deletion, while this biallelic marker may comprise about one to ten consecutive bases substituted, inserted or deleted.

20 Another aspect of the present invention is a method for determining the level of homology between a polypeptide code of SEQ ID Nos. 488 and 489 and a reference polypeptide sequence, comprising the steps of reading the polypeptide code of SEQ ID Nos. 488 and 489 and the reference polypeptide sequence through use of a computer program which determines homology levels and determining homology between the polypeptide code  
25 and the reference polypeptide sequence using the computer program.

Accordingly, another aspect of the present invention is a method for determining whether a nucleic acid code of the invention differs at one or more nucleotides from a reference nucleotide sequence comprising the steps of reading the nucleic acid code and the reference nucleotide sequence through use of a computer program which identifies  
30 differences between nucleic acid sequences and identifying differences between the nucleic acid code and the reference nucleotide sequence with the computer program. In some embodiments, the computer program is a program which identifies single nucleotide polymorphisms. The method may be implemented by the computer systems described above and the method illustrated in Figure 19. The method may also be performed by

reading at least 2, 5, 10, 15, 20, 25, 30, or 50 of the nucleic acid codes of the invention and the reference nucleotide sequences through the use of the computer program and identifying differences between the nucleic acid codes and the reference nucleotide sequences with the computer program. In other embodiments the computer based system may further comprise  
5 an identifier for identifying features within the nucleotide sequences of the nucleic acid codes of the invention or the amino acid sequences of the polypeptide codes of SEQ ID Nos. 488 and 489. An "identifier" refers to one or more programs which identifies certain features within the above-described nucleotide sequences of the nucleic acid codes of the invention or the amino acid sequences of the polypeptide codes of SEQ ID Nos. 488 and  
10 489. In one embodiment, the identifier may comprise a program which identifies an open reading frame in the cDNAs codes of SEQ ID Nos. 486 and 487.

Figure 20 is a flow diagram illustrating one embodiment of an identifier process 300 for detecting the presence of a feature in a sequence. The process 300 begins at a start state 302 and then moves to a state 304 wherein a first sequence that is to be checked for features  
15 is stored to a memory 115 in the computer system 100. The process 300 then moves to a state 306 wherein a database of sequence features is opened. Such a database would include a list of each feature's attributes along with the name of the feature. For example, a feature name could be "Initiation Codon" and the attribute would be "ATG." Another example would be the feature name "TAATAA Box" and the feature attribute would be "TAATAA".  
20 An example of such a database is produced by the University of Wisconsin Genetics Computer Group ([www.gcg.com](http://www.gcg.com)). Once the database of features is opened at the state 306, the process 300 moves to a state 308 wherein the first feature is read from the database. A comparison of the attribute of the first feature with the first sequence is then made at a state 310. A determination is then made at a decision state 316 whether the attribute of the  
25 feature was found in the first sequence. If the attribute was found, then the process 300 moves to a state 318 wherein the name of the found feature is displayed to the user. The process 300 then moves to a decision state 320 wherein a determination is made whether more features exist in the database. If no more features do exist, then the process 300 terminates at an end state 324. However, if more features do exist in the database, then the  
30 process 300 reads the next sequence feature at a state 326 and loops back to the state 310 wherein the attribute of the next feature is compared against the first sequence. It should be noted, that if the feature attribute is not found in the first sequence at the decision state 316, the process 300 moves directly to the decision state 320 in order to determine if any more features exist in the database. In another embodiment, the identifier may comprise a

molecular modeling program which determines the 3-dimensional structure of the polypeptides codes of SEQ ID Nos. 488 and 489. In some embodiments, the molecular modeling program identifies target sequences that are most compatible with profiles representing the structural environments of the residues in known three-dimensional protein structures. (See, e.g., Eisenberg et al., U.S. Patent No. 5,436,850 issued July 25, 1995). In another technique, the known three-dimensional structures of proteins in a given family are superimposed to define the structurally conserved regions in that family. This protein modeling technique also uses the known three-dimensional structure of a homologous protein to approximate the structure of the polypeptide codes of SEQ ID Nos. 488 and 489. (See e.g., Srinivasan, et al., U.S. Patent No. 5,557,535 issued September 17, 1996).

Conventional homology modeling techniques have been used routinely to build models of proteases and antibodies. (Sowdhamini et al., Protein Engineering 10:207, 215 (1997)). Comparative approaches can also be used to develop three-dimensional protein models when the protein of interest has poor sequence identity to template proteins. In some cases, proteins fold into similar three-dimensional structures despite having very weak sequence identities. For example, the three-dimensional structures of a number of helical cytokines fold in similar three-dimensional topology in spite of weak sequence homology. The recent development of threading methods now enables the identification of likely folding patterns in a number of situations where the structural relatedness between target and template(s) is not detectable at the sequence level. Hybrid methods, in which fold recognition is performed using Multiple Sequence Threading (MST), structural equivalencies are deduced from the threading output using a distance geometry program DRAGON to construct a low resolution model, and a full-atom representation is constructed using a molecular modeling package such as QUANTA.

According to this 3-step approach, candidate templates are first identified by using the novel fold recognition algorithm MST, which is capable of performing simultaneous threading of multiple aligned sequences onto one or more 3-D structures. In a second step, the structural equivalencies obtained from the MST output are converted into interresidue distance restraints and fed into the distance geometry program DRAGON, together with auxiliary information obtained from secondary structure predictions. The program combines the restraints in an unbiased manner and rapidly generates a large number of low resolution model confirmations. In a third step, these low resolution model confirmations are converted into full-atom models and subjected to energy minimization using the

molecular modeling package QUANTA. (See e.g., Aszódi et al., *Proteins: Structure, Function, and Genetics*, Supplement 1:38-42 (1997)).

The results of the molecular modeling analysis may then be used in rational drug design techniques to identify agents which modulate the activity of the polypeptide codes of SEQ ID Nos. 488 and 489. Accordingly, another aspect of the present invention is a method of identifying a feature within the nucleic acid codes of the invention or the polypeptide codes of SEQ ID Nos. 488 and 489 comprising reading the nucleic acid code(s) or the polypeptide code(s) through the use of a computer program which identifies features therein and identifying features within the nucleic acid code(s) or polypeptide code(s) with the computer program. In one embodiment, computer program comprises a computer program which identifies open reading frames. In a further embodiment, the computer program identifies structural motifs in a polypeptide sequence. In another embodiment, the computer program comprises a molecular modeling program. The method may be performed by reading a single sequence or at least 2, 5, 10, 15, 20, 25, 30, or 50 of the nucleic acid codes of the invention or the polypeptide codes of SEQ ID Nos. 488 and 489 through the use of the computer program and identifying features within the nucleic acid codes or polypeptide codes with the computer program. The nucleic acid codes of the invention or the polypeptide codes of SEQ ID Nos. 488 and 489 may be stored and manipulated in a variety of data processor programs in a variety of formats. For example, the nucleic acid codes of the invention or the polypeptide codes of SEQ ID Nos. 488 and 489 may be stored as text in a word processing file, such as MicrosoftWORD or WORDPERFECT or as an ASCII file in a variety of database programs familiar to those of skill in the art, such as DB2, SYBASE, or ORACLE. In addition, many computer programs and databases may be used as sequence comparers, identifiers, or sources of reference nucleotide or polypeptide sequences to be compared to the nucleic acid codes of the invention or the polypeptide codes of SEQ ID Nos. 488 and 489. The following list is intended not to limit the invention but to provide guidance to programs and databases which are useful with the nucleic acid codes of the invention or the polypeptide codes of SEQ ID No. 488 and 489. The programs and databases which may be used include, but are not limited to: MacPattern (EMBL), DiscoveryBase (Molecular Applications Group), GeneMine (Molecular Applications Group), Look (Molecular Applications Group), MacLook (Molecular Applications Group), BLAST and BLAST2 (NCBI), BLASTN and BLASTX (Altschul et al., *J. Mol. Biol.* 215: 403 (1990)), FASTA (Pearson and Lipman, *Proc. Natl. Acad. Sci. USA*, 85: 2444 (1988)), FASTDB (Brutlag et al. *Comp. App. Biosci.* 6:237-245. 1990), Catalyst (Molecular Simulations Inc.), Catalyst/SHAPE



(Molecular Simulations Inc.), Cerius<sup>2</sup>.DBAccess (Molecular Simulations Inc.), HypoGen (Molecular Simulations Inc.), Insight II, (Molecular Simulations Inc.), Discover (Molecular Simulations Inc.), CHARMM (Molecular Simulations Inc.), Felix (Molecular Simulations Inc.), DelPhi, (Molecular Simulations Inc.), QuanteMM, (Molecular Simulations Inc.),

5 Homology (Molecular Simulations Inc.), Modeler (Molecular Simulations Inc.), ISIS (Molecular Simulations Inc.), Quanta/Protein Design (Molecular Simulations Inc.), WebLab (Molecular Simulations Inc.), WebLab Diversity Explorer (Molecular Simulations Inc.), Gene Explorer (Molecular Simulations Inc.), SeqFold (Molecular Simulations Inc.), the EMBL/Swissprotein database, the MDL Available Chemicals Directory database, the MDL

10 Drug Data Report data base, the Comprehensive Medicinal Chemistry database, Derwent's World Drug Index database, the BioByteMasterFile database, the Genbank database, and the Genseqn database. Many other programs and data bases would be apparent to one of skill in the art given the present disclosure. Motifs which may be detected using the above programs include sequences encoding leucine zippers, helix-turn-helix motifs, glycosylation sites,

15 ubiquitination sites, alpha helices, and beta sheets, signal sequences encoding signal peptides which direct the secretion of the encoded proteins, sequences implicated in transcription regulation such as homeoboxes, acidic stretches, enzymatic active sites, substrate binding sites, and enzymatic cleavage sites.

It should be noted that the nucleic acid codes of the invention further encompass all

20 of the polynucleotides disclosed, described or claimed in the present invention. Also, it should be noted that the polypeptide codes of SEQ ID Nos. 488 and 489 further encompass all of the polypeptides disclosed, described or claimed in the present invention. Moreover, the present invention specifically contemplates the storage of such codes on computer readable media and computer systems individually or in combination, as well as the use of

25 such codes and combinations in the methods of section "VI. Computer-Related Embodiments."

### EXAMPLES

Several of the methods of the present invention are described in the following examples, which are offered by way of illustration and not by way of limitation. Many

30 other modifications and variations of the invention as herein set forth can be made without departing from the spirit and scope thereof and therefore only such limitations should be imposed as are indicated by the appended claims.

**Example 1****De Novo Identification Of Biallelic Markers**

The biallelic markers set forth in this application were isolated from human genomic sequences. To identify biallelic markers, genomic fragments were amplified, sequenced and  
5 compared in a plurality of individuals.

**DNA samples**

Donors were unrelated and healthy. They represented a sufficient diversity for being representative of a French heterogeneous population. The DNA from 100 individuals was extracted and tested for the *de novo* identification of biallelic markers.

10 DNA samples were prepared from peripheral venous blood as follows. Thirty ml of peripheral venous blood were taken from each donor in the presence of EDTA. Cells (pellet) were collected after centrifugation for 10 minutes at 2000 rpm. Red cells were lysed in a lysis solution (50 ml final volume: 10 mM Tris pH7.6; 5 mM MgCl<sub>2</sub>; 10 mM NaCl). The solution was centrifuged (10 minutes, 2000 rpm) as many times as necessary to  
15 eliminate the residual red cells present in the supernatant, after resuspension of the pellet in the lysis solution. The pellet of white cells was lysed overnight at 42°C with 3.7 ml of lysis solution composed of: (a) 3 ml TE 10-2 (Tris-HCl 10 mM, EDTA 2 mM) / NaCl 0.4 M; (b) 200 µl SDS 10%; and (c) 500 µl proteinase K (2 mg proteinase K in TE 10-2 / NaCl 0.4 M).

For the extraction of proteins, 1 ml saturated NaCl (6M) (1/3.5 v/v) was added.

20 After vigorous agitation, the solution was centrifuged for 20 minutes at 10000 rpm. For the precipitation of DNA, 2 to 3 volumes of 100% ethanol were added to the previous supernatant, and the solution was centrifuged for 30 minutes at 2000 rpm. The DNA solution was rinsed three times with 70% ethanol to eliminate salts, and centrifuged for 20 minutes at 2000 rpm. The pellet was dried at 37°C, and resuspended in 1 ml TE 10-1 or 1  
25 ml water. The DNA concentration was evaluated by measuring the OD at 260 nm (1 unit OD = 50 µg/ml DNA). To determine the presence of proteins in the DNA solution, the OD 260 / OD 280 ratio was determined. Only DNA preparations having a OD 260 / OD 280 ratio between 1.8 and 2 were used in the subsequent examples described below. DNA pools were constituted by mixing equivalent quantities of DNA from each individual.

**30 Amplification of genomic DNA by PCR**

Amplification of specific genomic sequences was carried out on pooled DNA samples obtained as described above.

**Amplification primers**

The primers used for the amplification of human genomic DNA fragments were defined with the OSP software (Hillier & Green, 1991). Preferably, primers included, upstream of the specific bases targeted for amplification, a common oligonucleotide tail useful for sequencing. Primers PU contain the following additional PU 5' sequence :

- 5 TGTAACACGACGGCCAGT; primers RP contain the following RP 5' sequence :  
CAGGAAACAGCTATGACC. Primers are listed in Figure 7.

#### Amplification

PCR assays were performed using the following protocol:

	Final volume	25 µl
10	DNA	2 ng/µl
	MgCl <sub>2</sub>	2 mM
	dNTP (each)	200 µM
	primer (each)	2.9 ng/µl
	Ampli Taq Gold DNA polymerase	0.05 unit/µl
15	PCR buffer (10x = 0.1 M TrisHCl pH8.3 0.5M KCl)	1x

DNA amplification was performed on a Genius II thermocycler. After heating at 94°C for 10 min, 40 cycles were performed. Cycling times and temperatures were: 30 sec at 94°C, 55°C for 1 min and 30 sec at 72°C. Holding for 7 min at 72°C allowed final elongation. The quantities of the amplification products obtained were determined on 96-  
20 well microtiter plates, using a fluorometer and Picogreen as intercalant agent (Molecular Probes).

#### **Sequencing of amplified genomic DNA and identification of biallelic polymorphisms**

Sequencing of the amplified DNA was carried out on ABI 377 sequencers. The sequences of the amplification products were determined using automated dideoxy  
25 terminator sequencing reactions with a dye terminator cycle sequencing protocol. The products of the sequencing reactions were run on sequencing gels and the sequences were determined using gel image analysis (ABI Prism DNA Sequencing Analysis software 2.1.2 version).

The sequence data were further evaluated to detect the presence of biallelic markers  
30 within the amplified fragments. The polymorphism search was based on the presence of superimposed peaks in the electrophoresis pattern resulting from different bases occurring at the same position. However, the presence of two peaks can be an artifact due to background noise. To exclude such an artifact, the two DNA strands were sequenced and a comparison between the two strands was carried out. In order to be registered as a polymorphic

sequence, the polymorphism had to be detected on both strands. Further, some biallelic single nucleotide polymorphisms were confirmed by microsequencing as described below.

Biallelic markers were identified in the analyzed fragments and are shown in Figure 1 and Table 2.

5

### Example 2

#### Genotyping Of Biallelic Markers

The biallelic markers identified as described above were further confirmed and their respective frequencies were determined through microsequencing. Microsequencing was carried out on individual DNA samples obtained as described herein.

#### 10 Microsequencing primers

Amplification of genomic DNA fragments from individual DNA samples was performed as described in Example 1 using the same set of PCR primers (Figure 7). Microsequencing was carried out on the amplified fragments using specific primers. See Figure 6. The preferred primers used in microsequencing had about 19 nucleotides in length and hybridized just upstream of the considered polymorphic base.

The microsequencing reactions were performed as follows: 5 µl of PCR products were added to 5 µl purification mix (2U SAP (Shrimp alkaline phosphate) (Amersham E70092X)); 2U Exonuclease I (Amersham E70073Z); and 1 µl SAP buffer (200 mM Tris-HCl pH8, 100 mM MgCl<sub>2</sub>) in a microtiter plate. The reaction mixture was incubated 20 minutes at 37°C, and denatured 10 minutes at 94°C afterwards. To each well was then added 20 µl of microsequencing reaction mixture containing: 10 pmol microsequencing oligonucleotide (19mers, GENSET, crude synthesis, 5 OD), 1 U Thermosequenase (Amersham E79000G), 1.25 µl Thermosequenase buffer (260 mM Tris HCl pH 9.5, 65 mM MgCl<sub>2</sub>), and the two appropriate fluorescent ddNTPs complementary to the nucleotides at 25 the polymorphic site corresponding to both polymorphic bases (11.25 nM TAMRA-ddTTP ; 16.25 nM ROX-ddCTP ; 1.675 nM REG-ddATP ; 1.25 nM RHO-ddGTP ; Perkin Elmer, Dye Terminator Set 401095). After 4 minutes at 94°C, 20 PCR cycles of 15 sec at 55°C, 5 sec at 72°C, and 10 sec at 94°C were carried out in a Tetrad PTC-225 thermocycler (MJ Research). The microtiter plate was centrifuged 10 sec at 1500 rpm. The unincorporated 30 dye terminators were removed by precipitation with 19 µl MgCl<sub>2</sub> 2mM and 55 µl 100 % ethanol. After 15 minute incubation at room temperature, the microtiter plate was centrifuged at 3300 rpm 15 minutes at 4°C. After discarding the supernatants, the microplate was evaporated to dryness under reduced pressure (Speed Vac). Samples were resuspended in 2.5 µl formamide EDTA loading buffer and heated for 2 min at 95°C. 0.8 µl

microsequencing reaction were loaded on a 10 % (19:1) polyacrylamide sequencing gel. The data were collected by an ABI PRISM 377 DNA sequencer and processed using the GENESCAN software (Perkin Elmer).

#### **Frequency of biallelic markers**

- 5       Frequencies are reported for the less common allele only and are shown in Figure 1. The frequencies for both alleles are shown for the MGST-II, ME1, UGT1A7 and UGT2B4 genes in Figures 9, 10, 11, and 12, respectively.

#### **Example 3**

##### **Association Between Asthma and the Biallelic Markers of the MGST-II Gene**

#### **10   Collection of DNA samples from trait positive and control individuals**

- The disease trait followed in this association study was asthma, a disease involving the leukotriene pathway. The asthmatic population corresponded to 298 individuals that took part in a clinical study for the evaluation of the anti-asthmatic drug zileuton. More than 90 % of these 297 asthmatic individuals had a Caucasian ethnic background. The  
15   control population corresponded to 178 individuals from a random US Caucasian population.

#### **Genotyping of affected and control individuals**

- The general strategy to perform the association studies was to individually scan the DNA samples from all individuals in each of the populations described above in order to  
20   establish the allele frequencies of the above described biallelic markers in each of these populations.

- Allelic frequencies of the above-described biallelic marker alleles in each population were determined by performing microsequencing reactions on amplified fragments obtained by genomic PCR performed on the DNA samples from each individual. Genomic PCR and  
25   microsequencing were performed as detailed above in Examples 1 and 2 using the described PCR and microsequencing primers.

#### **Haplotype frequency analysis**

- None of the single marker alleles showed a significant association with asthma however, significant results were obtained in haplotype studies. Allelic frequencies were  
30   useful to check that the markers used in the haplotype studies meet the Hardy-Weinberg proportions (random mating).

The results of the haplotype analysis using 13 biallelic markers (12-421-135, 12-421-140, 12-430-80, 12-441-233, 12-442-133, 12-447-58, 12-455-326, 12-461-299, 12-453-429, 12-424-198, 12-454-363, 12-458-196 and 12-426-154) are shown in Figure 13.

Haplotype analysis for association of MGST-II biallelic markers and asthma was performed by estimating the frequencies of all possible 2, 3 and 4 marker haplotypes in the asthmatic and control populations described above. Haplotype estimations were performed by applying the Expectation-Maximization (EM) algorithm (Excoffier and Slatkin, *Mol. Biol. Evol.*, 12:921-927, 1995), using the EM-HAPLO program (Hawley et al., *Am. J. Phys. Anthropol.*, 18:104, 1994) as described above. Estimated haplotype frequencies in the asthmatic and control population were compared by means of a chi-square statistical test (one degree of freedom).

Figure 13 shows the most significant haplotypes obtained. Haplotype No. 6 consisting of three biallelic markers (12-455-326, 12-453-429 and 12-424-198) had a p-value of  $3.2 \times 10^{-5}$  and an odds ratio of 12.22. Estimated haplotype frequencies were 11.8 % in the cases and 1.1 % in the controls. Haplotype No. 18 consisting of four biallelic markers (12-455-326, 12-453-429, 12-424-198 and 12-454-363) had a p-value of  $1.2 \times 10^{-6}$  and an odds ratio of 100.00. Both haplotypes are related as three out of four biallelic marker alleles (G at 12-455-326, C at 12-453-429 and T and 12-454-363) are common to both haplotypes. Haplotype No. 19 consisting of four biallelic markers (12-447-58, 12-455-326, 12-461-299 and 12-453-429) had a p-value of  $8.2 \times 10^{-6}$  and an odds ratio of 100.00. Markers 12-455-326 and 12-453-429 are common in all three significant haplotypes; therefore, they represent preferred markers for the diagnosis of asthma. Haplotypes Nos. 6, 18 and 19 are strongly associated with asthma. Haplotypes Nos. 7-17 and 20-30 also showed very significant association (see Figure 13).

The statistical significance of the results obtained for the haplotype analysis was evaluated by a phenotypic permutation test reiterated 1000 or 10,000. For this computer simulation, data from the asthmatic and control individuals were pooled and randomly allocated to two groups which contained the same number of individuals as the case-control populations used to produce the data summarized in Figure 13. A haplotype analysis was then run on these artificial groups for the 3 markers included in haplotype No. 6 (haplotype GCT), the 4 markers included in haplotype No. 18 (haplotype GCTG) and the 4 markers included in haplotype No. 19 (haplotype CATT), all of which showed a strong association with asthma. This experiment was reiterated 1000 and 10,000 times and the results are shown in Figure 14. These results demonstrate that among 1000 iterations only 3 and among 10,000 iterations only 31 of the obtained haplotypes had a p-value comparable to the one obtained for haplotype No. 6 (haplotype GCT). These results demonstrate that among 1000 iterations 0 and among 10,000 iterations only 5 of the obtained haplotypes had a p-

value comparable to the one obtained for haplotype No. 18 (haplotype GCTG). These results further demonstrate that among 1000 iterations only 12 and among 10,000 iterations only 76 of the obtained haplotypes had a p-value comparable to the one obtained for haplotype No. 19 (haplotype CATT). These results clearly validate the statistical significance of the association between the haplotypes shown in Figure 13 and asthma.

#### **Example 4**

##### **Association Between Side Effects Upon Treatment With the Anti-Asthmatic Drug Zileuton (Zyflo™) and the Biallelic Markers of the MGST-II Gene**

###### **Collection of DNA samples from trait positive and control individuals**

10 The side effect examined in this study was the hepatotoxicity experienced by asthmatic individuals as a result of their treatment with Zileuton as part of a clinical study. Asthmatic individuals were unrelated and more than 90% of the individuals had a Caucasian ethnic background. Hepatotoxicity was monitored by measuring the serum levels of alanine aminotransferase (ALT), which is a sensitive indicator of liver cell damage.

15 More than 90% of the asthmatic individuals participating in this study did not experience Zileuton-associated ALT increase compared to their ALT levels prior to zileuton intake. As mentioned above, an association study is more informative if the populations considered present extreme phenotypes. Therefore, the asthmatic individuals, which were selected for the side effect positive trait (ALT+), corresponded to 89 individuals that  
20 presented at least 3 times the upper limit of normal (ULN) level of ALT. On the other side, the asthmatic individuals that were selected for the side effect negative trait (ALT-) corresponded to 208 individuals that presented less than 1xULN of ALT. ALT+ and ALT- populations corresponded to 4% and 35% respectively of the total asthmatic individuals that participated in this study.

###### **25 Genotyping of affected and control individuals**

The general strategy to perform the Association studies was to individually scan the DNA samples from all individuals in each of the populations described above in order to establish the allele frequencies of the above described biallelic markers in each of these populations.

30 Allelic frequencies of the above-described biallelic marker alleles in each population were determined by performing microsequencing reactions on amplified fragments obtained by genomic PCR performed on the DNA samples from each individual. Genomic PCR and microsequencing were performed as detailed above in Examples 1 and 2 using the described PCR and microsequencing primers.

### Haplotype frequency analysis

None of the single marker alleles showed a significant association with hepatotoxicity to zileuton however, significant results were obtained in haplotype studies.

The results of the haplotype analysis using 13 biallelic markers (12-421-135, 12-421-140, 12-430-80, 12-441-233, 12-442-133, 12-447-58, 12-455-326, 12-461-299, 12-453-429, 12-424-198, 12-454-363, 12-458-196 and 12-426-154) are shown in Figure 15.

Haplotype analysis for association of MGST-II biallelic markers and asthma was performed by estimating the frequencies of all possible 2, 3 and 4 marker haplotypes in the ALT+ and ALT- populations described above. Haplotype estimations were performed by applying the Expectation-Maximization (EM) algorithm (Excoffier and Slatkin, *Mol. Biol. Evol.*, 12:921-927, 1995), using the EM-HAPLO program (Hawley et al., *Am. J. Phys. Anthropol.*, 18:104, 1994) as described above. Estimated haplotype frequencies in the ALT+ and ALT- populations were compared by means of a chi-square statistical test (one degree of freedom).

Figure 15 shows the most significant haplotypes obtained. Haplotype No. 6 consisting of three biallelic markers (12-441-233, 12-461-299 and 12-453-429) presented a p-value of  $1.5 \cdot 10^{-5}$  and an odd-ratio of 3.63. Estimated haplotype frequencies were 15.7 % in the cases and 4.9 % in the controls. Haplotype No. 19 consisting of four biallelic markers (12-441-233, 12-461-299, 12-453-429 and 12-426-154) had a p-value of  $5.2 \cdot 10^{-7}$  and an odd ratio of 5.75. Estimated haplotype frequencies were 14.1 % in the cases and 2.8 % in the controls. Both haplotypes showed strong association with elevated serum ALT level upon treatment with zileuton. Both haplotypes are related as three out of four biallelic marker alleles (C at 12-441-233, T at 12-461-299 and T at 12-453-429) are common to both haplotypes. Haplotypes Nos. 7-18 and 20-31 of Figure 15 also showed very significant association.

The statistical significance of the results obtained for the haplotype analysis was evaluated by a phenotypic permutation test reiterated 1000 or 10,000 times on a computer. For this computer simulation, data from the ALT+ and ALT- populations were pooled and randomly allocated to two groups which contained the same number of individuals as the ALT+ and ALT- populations used to produce the data summarized in Figure 15. A haplotype analysis was then run on these artificial groups for the 3 markers included in haplotype No. 6 (haplotype CTT) and for the 4 markers included in haplotype No. 19 (haplotype CTTA) which, showed the strongest association with secondary effects to zileuton. This experiment was reiterated 1000 and 10,000 times and the results are shown in



Figure 16. These results demonstrate that among 1000 iterations only 1 and among 10,000 iterations only 12 of the obtained haplotypes had a p-value comparable to the one obtained for haplotype No. 6 (haplotype CTT). These results further demonstrate that among 1000 iterations only 1 and among 10,000 iterations only 7 of the obtained haplotypes had a p-value comparable to the one obtained for haplotype No. 19 (haplotype CTTA). These results clearly validate the statistical significance of the association between hepatotoxicity to Zylflo™ and the haplotypes shown in Figure 15.

#### **Example 5**

##### **Identification of Mutations and of Low Frequency Alleles of the MGST-II Gene**

Exons 3-5, the 5'UTR region and the 3'region of the MGST-II gene were screened for mutations by comparing their sequence in individuals exhibiting elevated ALT levels upon treatment with zileuton (ALT+) and in individuals showing normal ALT levels upon treatment with zileuton (ALT-). ALT + and ALT- individuals are further described in Example 4. Intron sequences immediately flanking the exons were also screened.

To identify mutations, fragments of the MGST-II gene were amplified, sequenced and compared in ALT+ and ALT- individuals. DNA samples from each individual were processed separately.

##### **DNA samples**

Individual DNA samples were obtained as described in Example 1.

##### **Amplification of the MGST-II gene**

Amplification primers are described in Table 5 and PCR assays were performed as described in Example 1.

##### **Sequencing of amplified genomic DNA: identification of mutations and of low frequency polymorphisms**

Sequencing of the amplified DNA was carried out on ABI 377 sequencers. The sequences of the amplification products were determined using automated dideoxy terminator sequencing reactions with a dye terminator cycle sequencing protocol. The products of the sequencing reactions were run on sequencing gels and the sequences were determined using gel image analysis (ABI Prism DNA Sequencing Analysis software 2.1.2 version).

The sequence data was further analyzed to detect the presence of mutations and of low frequency alleles. The sequences obtained with 79 ALT+ individuals and 105 ALT- individuals were compared. New polymorphisms/mutations were detected and the genotype of each individual for these markers was determined. Results are shown below:

Marker ID	Position in MGST-II gene	Least common allele/ mutation	Original allele	Genotype of ALT+ and ALT- individuals
10-286-289	5'UTR	G	C	79 ALT+ C/C 104 ALT- C/C 1 ALT- GC
10-286-345	5'UTR	T	A	65 ALT+ A/A 12 ALT+ A/T 2 ALT+ T/T 82 ALT- A/A 19 ALT- A/T 3 ALT- T/T
10-286-375	5'UTR	G	A	78 ALT+ A/A 1 ALT+ A/G 104 ALT- A/A
10-523-232	Exon 3	T	C	75 ALT+ C/C 100 ALT- C/C 1 ALT- C/T
10-289-201	Exon 4	C	T	76 ALT+ T/T 2 ALT+ C/T 100 ALT- T/T 1 ALT- C/T
10-290-37	Exon 5	T	C	78 ALT+ C/C 1 ALT+ C/T 104 ALT- C/C
10-290-326	3' region	A	G	79 ALT+ G/G 103 ALT- G/G 1 ALT- A/G
10-290-328	3' region	deletion	-	1 ALT+ deletion 78 ALT+ - 104 ALT- -

Eight polymorphisms were identified in the region of the MGST-II gene that was scanned. Three mutations were identified in the 3'UTR region. One mutation in exon 4 causes an amino acid substitution (Tyr→His) at the polypeptide level. A mutation in exon 5 introduces a stop codon into the ORF leading to the expression of a truncated MGST-II polypeptide. These mutations modify the specificity, activity and function of the MGST-II enzyme and therefore represent functional mutations of the MGST-II gene.

#### Example 6

##### Preparation of Antibody Compositions to MGST-II Variants

10 Preferably antibody compositions, specifically binding the 93-His variant or the SEQ ID No. 489 variant of MGST-II, are prepared.

Substantially pure protein or polypeptide is isolated from transfected or transformed cells containing an expression vector encoding the MGST-II protein or a portion thereof. The concentration of protein in the final preparation is adjusted, for example, by concentration on

an Amicon filter device, to the level of a few micrograms per ml. Monoclonal or polyclonal antibodies to the protein can then be prepared as follows:

#### **Monoclonal Antibody Production by Hybridoma Fusion**

Monoclonal antibody to epitopes in the MGST-II protein or a portion thereof can be  
5 prepared from murine hybridomas according to the classical method of Kohler and Milstein (*Nature*, 256:495, 1975) or derivative methods thereof (see Harlow and Lane, *Antibodies A Laboratory Manual*, Cold Spring Harbor Laboratory, pp. 53-242, 1988).

Briefly, a mouse is repetitively inoculated with a few micrograms of the MGST-II protein or a portion thereof over a period of a few weeks. The mouse is then sacrificed, and the  
10 antibody producing cells of the spleen isolated. The spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by  
15 detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as ELISA, as originally described by Engvall, E., *Meth. Enzymol.* 70:419 (1980), and derivative methods thereof. Selected positive clones can be expanded and their monoclonal antibody product harvested for use. Detailed procedures for monoclonal antibody production are described in Davis, L. et al. *Basic Methods in Molecular Biology* Elsevier, New York. Section  
20 21-2.

#### **Polyclonal Antibody Production by Immunization**

Polyclonal antiserum containing antibodies to heterogeneous epitopes in the MGST-II protein or a portion thereof can be prepared by immunizing suitable non-human animal with the MGST-II protein or a portion thereof, which can be unmodified or modified to enhance  
25 immunogenicity. A suitable non-human animal is preferably a non-human mammal is selected, usually a mouse, rat, rabbit, goat, or horse. Alternatively, a crude preparation which, has been enriched for MGST-II concentration can be used to generate antibodies. Such proteins, fragments or preparations are introduced into the non-human mammal in the presence of an appropriate adjuvant (e.g. aluminum hydroxide, RIBI, etc.) which is known  
30 in the art. In addition the protein, fragment or preparation can be pretreated with an agent which will increase antigenicity, such agents are known in the art and include, for example, methylated bovine serum albumin (mBSA), bovine serum albumin (BSA), Hepatitis B surface antigen, and keyhole limpet hemocyanin (KLH). Serum from the immunized animal is collected, treated and tested according to known procedures. If the serum contains

polyclonal antibodies to undesired epitopes, the polyclonal antibodies can be purified by immunoaffinity chromatography.

Effective polyclonal antibody production is affected by many factors related both to the antigen and the host species. Also, host animals vary in response to site of inoculations and dose, with both inadequate or excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. Techniques for producing and processing polyclonal antisera are known in the art, see for example, Mayer and Walker (1987). An effective immunization protocol for rabbits can be found in Vaitukaitis, J. et al. *J. Clin. Endocrinol. Metab.* 33:988-991 (1971).

Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against known concentrations of the antigen, begins to fall. (See, for example, Ouchterlony, O. et al., Chap. 19 in: *Handbook of Experimental Immunology* D. Wier (ed) Blackwell, 1973). Plateau concentration of antibody is usually in the range of 0.1 to 0.2 mg/ml of serum. Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as described, for example, by Fisher, D., (Chap. 42: *Manual of Clinical Immunology*, 2d Ed. Rose and Friedman, Eds., *Amer. Soc. For Microbiol.*, Washington, D.C., 1980).

Antibody preparations prepared according to either the monoclonal or the polyclonal protocol are useful in quantitative immunoassays which determine concentrations of antigen-bearing substances in biological samples; they are also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample. The antibodies may also be used in therapeutic compositions for killing cells expressing the protein or reducing the levels of the protein in the body.

It should be noted that in the accompanying Sequence Listing, all instances of the symbol "n" in the nucleic acid sequences mean that the nucleotide can be adenine, guanine, cytosine or thymine.

In some instances, the polymorphic bases of the biallelic markers alter the identity of amino acids in the encoded polypeptide. This is indicated in the accompanying Sequence Listing by use of the feature VARIANT, placement of a Xaa at the position of the polymorphic amino acid, and definition of Xaa as the two alternative amino acids. For example, if one allele of a biallelic marker is the codon CAC, which encodes histidine, while the other allele of the biallelic marker is CAA, which encodes glutamine, the

145

Sequence Listing for the encoded polypeptide will contain an Xaa at the location of the polymorphic amino acid. In this instance, Xaa would be defined as being histidine or glutamine.

- In other instances, Xaa may indicate an amino acid whose identity is unknown
- 5 because of nucleotide sequence ambiguity. In this instance, the feature UNSURE is used, Xaa is placed at the position of the unknown amino acid, and Xaa is defined as being any of the 20 amino acids or a limited number of amino acids suggested by the genetic code.

SEQUENCE LISTING FREE TEXT

The following free text appears in the accompanying sequence listing:

- biallelic
- marker
- 5 drug metabolism
- Homo Sapiens
- allele
- polymorphic base
- misc\_binding
- 10 potential
- complement
- potential complement
- primer\_bind
- upstream amplification primer
- 15 upstream amplification primer, complement
- downstream amplification primer
- potential probe
- misc\_feature
- 5' regulatory region
- 20 exon
- 5'UTR
- CDS
- 3'UTR
- polyA\_signal
- 25 PRT
- VARIANT
- Stop
- Artificial Sequence
- sequencing oligonucleotide PrimerPU
- 30 sequencing oligonucleotide PrimerRP

WHAT IS CLAIMED IS:

1. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of the sequences  
5 described in Figure 2 and the complements thereof.
2. A polynucleotide according to claim 1, wherein said span includes a DME-related biallelic marker in said sequence.
- 10 3. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of the sequences described in Figure 3 and the complements thereof, wherein said span includes a DME-related biallelic marker in said sequence with the alternative allele present at said biallelic marker.  
15
4. An isolated, purified or recombinant polynucleotide consisting essentially of a contiguous span of 8 to 50 nucleotides of a sequence selected from the group consisting of the sequences described in Figure 3 and the complements thereof, wherein said span includes a DME-related biallelic marker in said sequence with the original allele present at  
20 said biallelic marker.
5. An isolated, purified or recombinant polynucleotide consisting essentially of a contiguous span of 8 to 50 nucleotides of a sequence selected from the group consisting of the sequences described in Figure 4 and the complements thereof, wherein said span  
25 includes a DME-related biallelic marker in said sequence.
6. A polynucleotide according to any one of claims 2 to 5, wherein said contiguous span is 18 to 35 nucleotides in length and said biallelic marker is within 4 nucleotides of the center of said polynucleotide.  
30
7. A polynucleotide according to claim 6, wherein said polynucleotide consists of said contiguous span and said contiguous span is 25 nucleotides in length and said biallelic marker is at the center of said polynucleotide.

148

8. A polynucleotide according to claim 1, wherein the 3' end of said contiguous span is present at the 3' end of said polynucleotide.
9. A polynucleotide according to any one of claims 2 to 5, wherein the 3' end of said  
5 contiguous span is located at the 3' end of said polynucleotide and said biallelic marker is present at the 3' end of said polynucleotide.
10. A polynucleotide according to claim 8, wherein the 3' end of said polynucleotide is located within 20 nucleotides upstream of a DME-related biallelic marker in said sequence.  
10
11. An isolated, purified or recombinant polynucleotide consisting essentially of a contiguous span of 8 to 50 nucleotides in a sequence selected from the group consisting of the sequences described in Figure 3, the sequences described in Figure 4, and the complements thereof, wherein the 3' end of said contiguous span is located at the 3' end of  
15 said polynucleotide, and wherein the 3' end of said polynucleotide is located within 20 nucleotides upstream of a DME-related biallelic marker in said sequence.
12. A polynucleotide according to either claim 10 or 11, wherein the 3' end of said polynucleotide is located 1 nucleotide upstream of a DME-related biallelic marker in said  
20 sequence.
13. A polynucleotide according to claim 1, wherein said polynucleotide consists essentially of a sequence selected from the sequences described in Figure 6.
- 25 14. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of a sequence selected from the sequences described in Figure 5.
15. A polynucleotide consisting essentially of a sequence selected from the sequences described in Figure 7.  
30
16. A polynucleotide consisting essentially of a sequence selected from the sequences described in Figure 8.



17. A polynucleotide according to any one of claims 1, 3, 4, 5, 11, 14, 15 and 16 wherein said contiguous span comprises at least 15 contiguous nucleotides in said sequence.
18. A polynucleotide according to any one of claims 1, 3, 4, 5, 11, 14, 15 and 16 wherein  
5 said contiguous span comprises at least 20 contiguous nucleotides in said sequence.
19. A polynucleotide according to any one of claims 1, 3, 4, 5, 11, 14, 15 and 16 wherein said contiguous span comprises at least 25 contiguous nucleotides in said sequence.
- 10 20. A polynucleotide according to any one of claims 1, 3, 4, 5, 11, 14, 15 and 16 attached to a solid support.
21. An array of polynucleotides comprising at least one polynucleotide according to claim 20.
- 15 22. An array according to claim 21, wherein said array is addressable.
23. A polynucleotide according to any one of claims 1, 3, 4, 5, 11, 14, 15 and 16, further comprising a label.
- 20 24. A method of genotyping comprising determining the identity of a nucleotide at a DME-related biallelic marker or MGST-II-related biallelic marker in a biological sample.
- 25 25. A method according to claim 24, wherein said DME-related biallelic marker is selected from the biallelic markers described in Figure 1.
26. A method according to claim 25, wherein said DME-related biallelic marker is selected from the group consisting of the biallelic markers found in Figures 9, 10, 11 and 12.
- 30 27. A method according to claim 25, wherein said DME-related biallelic marker is selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284.

28. A method according to claim 24, wherein said biological sample is derived from a single subject.
- 5 29. A method according to claim 28, wherein the identity of the nucleotides at said biallelic marker is determined for both copies of said biallelic marker present in said subject's genome.
30. A method according claim 24, wherein said biological sample is derived from  
10 multiple subjects.
31. A method according to claim 24, further comprising amplifying a portion of said sequence comprising the biallelic marker prior to said determining step.
- 15 32. A method according to claim 31, wherein said amplifying is performed by PCR.
33. A method according to claim 24, wherein said determining is performed by a hybridization assay.
- 20 34. A method according to claim 24, wherein said determining is performed by a sequencing assay.
35. A method according to claim 24, wherein said determining is performed by a microsequencing assay.
- 25 36. A method according to claim 24, wherein said determining is performed by an enzyme-based mismatch detection assay.
37. A method of determining the frequency in a population of an allele of a DME-  
30 related biallelic marker or MGST-II-related biallelic marker, comprising:  
a) genotyping individuals from said population for said biallelic marker according to the method of claim 24; and  
b) determining the proportional representation of said biallelic marker in said population.

38. A method according to claim 37, wherein said DME-related biallelic marker is selected from the biallelic markers described in Figure 1.
- 5 39. A method according to claim 38, wherein said DME-related biallelic marker is selected from the group consisting of the biallelic markers found in Figures 9, 10, 11 and 12.
40. A method according to claim 38, wherein said DME-related biallelic marker is selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233,  
10 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284.
41. A method according to claim 37, wherein said genotyping of step a) is performed on  
15 each individual of said population.
42. A method according to claim 37, wherein said genotyping is performed on a single biological sample derived from said population.
- 20 43. A method of detecting an association between an allele and a phenotype, comprising the steps of:
- a) determining the frequency of at least one DME-related biallelic marker or MGST-II-related biallelic marker allele in a trait positive population according to the method of claim 37;
  - 25 b) determining the frequency of said DME-related biallelic marker or MGST-II-related biallelic marker allele in a control population according to the method of claim 37; and
  - c) determining whether a statistically significant association exists between said allele and said phenotype.
- 30 44. A method of estimating the frequency of a haplotype for a set of biallelic markers in a population, comprising:
- a) genotyping each individual in said population for at least one DME-related biallelic marker or MGST-II-related biallelic marker according to claim 24;

152

b) genotyping each individual in said population for a second biallelic marker by determining the identity of the nucleotides at said second biallelic marker for both copies of said second biallelic marker present in the genome; and

5 c) applying a haplotype determination method to the identities of the nucleotides determined in steps a) and b) to obtain an estimate of said frequency.

45. A method according to claim 44, wherein said haplotype determination method is selected from the group consisting of asymmetric PCR amplification, double PCR amplification of specific alleles, the Clark method, and an expectation maximization  
10 algorithm.

46. A method according to claim 44, wherein said DME-related biallelic marker is selected from the biallelic markers described in Figure 1.

15 47. A method according to claim 46, wherein said DME-related biallelic marker is selected from the group consisting of the biallelic markers found in Figures 9, 10, 11 and 12.

48. A method according to claim 46, wherein said DME-related biallelic marker is selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233,  
20 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284.

49. A method according to claim 46, wherein said haplotype comprises one of the  
25 following sets of biallelic markers:

12-455-326, 12-453-429 and 12-424-198;

12-455-326, 12-453-429, 12-424-198 and 12-454-363;

12-447-58, 12-455-326, 12-461-299 and 12-453-429;

12-441-233, 12-461-299 and 12-453-429;

30 12-441-233, 12-461-299, 12-453-429 and 12-426-154;

12-426-154, and 12-424-198;

12-426-154, 12-461-299, and 12-424-198;

10-428-219, and 12-724-225;

12-128-225, 12-156-91, 12-139-380, and 12-140-134;

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153

12-148-311, 12-156-91, 12-139-380, 12-140-134;  
10-470-25, and 12-652-203; and  
10-470-25, 12-637-219, and 12-652-203.

- 5 50. A method according to claim 46, wherein said haplotype comprises one of the following sets of biallelic markers:

12-455-326, 12-453-429 and 12-424-198;  
12-455-326, 12-453-429, 12-424-198 and 12-454-363;  
12-447-58, 12-455-326, 12-461-299 and 12-453-429;  
10 12-426-154, and 12-424-198;  
10-428-219, and 12-724-225;  
12-128-225, 12-156-91, 12-139-380, and 12-140-134; and  
10-470-25, and 12-652-203.

- 15 51. A method of detecting an association between a haplotype and a phenotype, comprising the steps of:

a) estimating the frequency of at least one haplotype in a trait positive population according to the method of claim 44;  
b) estimating the frequency of said haplotype in a control population  
20 according to the method of claim 44; and  
c) determining whether a statistically significant association exists between said haplotype and said phenotype.

52. A method according to either claim 43 or 51, wherein said control population is a  
25 trait negative population.

53. A method according to either claim 43 or 51, wherein said case control population is a random population.

- 30 54. A method according to claim 51, wherein said haplotype is selected from the group of DME-related biallelic marker consisting of the biallelic markers found in Figures 9, 10, 11 and 12.

55. A method according to claim 51, wherein said haplotype is selected from the group of DME-related biallelic marker consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-  
5 219, 12-721-440, and 10-420-284.

56. A method according to claim 51, wherein said haplotype comprises one of the following sets of DME-related biallelic preferred markers:

12-455-326, 12-453-429 and 12-424-198;  
10 12-455-326, 12-453-429, 12-424-198 and 12-454-363;  
12-447-58, 12-455-326, 12-461-299 and 12-453-429;  
12-441-233, 12-461-299 and 12-453-429;  
12-441-233, 12-461-299, 12-453-429 and 12-426-154;  
12-426-154, and 12-424-198;  
15 12-426-154, 12-461-299, and 12-424-198;  
10-428-219, and 12-724-225;  
12-128-225, 12-156-91, 12-139-380, and 12-140-134;  
12-148-311, 12-156-91, 12-139-380, 12-140-134;  
10-470-25, and 12-652-203; and  
20 10-470-25, 12-637-219, and 12-652-203.

57. A method according to claim 51, wherein said haplotype comprises one of the following sets of DME-related biallelic markers:

12-455-326, 12-453-429 and 12-424-198;  
25 12-455-326, 12-453-429, 12-424-198 and 12-454-363;  
12-447-58, 12-455-326, 12-461-299 and 12-453-429;  
12-426-154, and 12-424-198;  
10-428-219, and 12-724-225;  
12-128-225, 12-156-91, 12-139-380, and 12-140-134; and  
30 10-470-25, and 12-652-203.

58. A method according to claim 43, wherein each of said determining of steps a) and b) is performed on a single pooled biological sample derived from each of said populations.

59. A method according to claim 43, wherein said genotyping of steps a) and b) is performed separately on biological samples derived from each individual in said populations.
- 5 60. A method according to either claim 43 or 51, wherein said phenotype is a response to a drug.
61. A method according to either claim 43 or 51, wherein said phenotype is a side effect to a drug.
- 10 62. A method according to either claim 43 or 51, wherein said phenotype is a disease involving the metabolic conversion of xenobiotics.
63. A method according to claim 43, wherein the identity of the nucleotides at all of the  
15 biallelic markers described in Figure 1 is determined in steps a) and b).
64. A computer readable medium having stored thereon a sequence selected from the group consisting of a nucleic acid code comprising a contiguous span of at least 12 nucleotides of a sequence described in Figure 2, Figure 3, Figure 5 and the complements  
20 thereof; wherein said contiguous span of a sequence described in Figure 3 comprises a DME-related biallelic marker with the alternative allele present at said biallelic marker.
65. A computer system comprising a processor and a data storage device wherein said data storage device has stored thereon a sequence selected from the group consisting of a  
25 nucleic acid code comprising a contiguous span of at least 12 nucleotides of a sequence described in Figure 2, Figure 3, Figure 5 and the complements thereof; wherein said contiguous span of a sequence described in Figure 3 comprises a DME-related biallelic marker with the alternative allele present at said biallelic marker.
- 30 66. The computer system of Claim 53 further comprising a sequence comparer and a data storage device having reference sequences stored thereon.
67. The computer system of Claim 54 wherein said sequence comparer comprises a computer program which indicates polymorphisms.

68. A method for comparing a first sequence to a reference sequence, comprising the steps of:

5 a) reading said first sequence and said reference sequence through use of a computer program which compares sequences; and

b) determining differences between said first sequence and said reference sequence with said computer program;

wherein said first sequence is selected from the group consisting of a nucleic acid comprising a contiguous span of at least 12 nucleotides of a sequence described in Figure 2, Figure 3, Figure 5 and the complements thereof; wherein said contiguous span of a sequence described in Figure 3 comprises a DME-related biallelic marker with the alternative allele present at said biallelic marker.

69. The method of Claim 68, wherein said step b) comprises identifying polymorphisms.

15

70. A method of administering a drug or treatment comprising:

a) obtaining a nucleic acid sample from an individual;

20 b) determining the identity of the polymorphic base of at least one DME-related biallelic marker or MGST-II-related biallelic marker according to the method of claim 24 which is associated with a positive response to said drug or treatment, or at least one DME-related biallelic marker or MGST-II-related biallelic marker or which is associated with a negative response to said drug or treatment; and

25 c) administering said drug or treatment to said individual if said nucleic acid sample contains at least one biallelic marker associated with a positive response to said drug or treatment, or if said nucleic acid sample lacks at least one biallelic marker associated with a negative response to said drug or treatment.

71. A method of selecting an individual for inclusion in a clinical trial of a drug or treatment comprising:

30 a) obtaining a nucleic acid sample from an individual;

b) determining the identity of the polymorphic base of at least one DME-related biallelic marker or MGST-II-related biallelic marker according to the method of claim 24 which is associated with a positive response to said drug or treatment, or



157

at least one biallelic marker associated with a negative response to said drug or treatment in said nucleic acid sample; and

5 c) including said individual in said clinical trial if said nucleic acid sample contains at least one biallelic marker which is associated with a positive response to said drug or treatment, or if said nucleic acid sample lacks at least one biallelic marker associated with a negative response to said drug or treatment.

72. A method according to claim 70, wherein said administering step comprises administering said drug or treatment to said individual if said nucleic acid sample contains  
10 at least one biallelic marker associated with a positive response to said drug treatment, and said nucleic acid sample lacks at least one biallelic marker associated with a negative response to said drug or treatment.

73. The method according to either claim 70 or 71, wherein said DME-related biallelic  
15 marker is selected from the group consisting of the biallelic markers found in Figure 1.

74. A method according to claim 73, wherein said DME-related biallelic marker is selected from the group consisting of the biallelic markers found in Figures 9, 10, 11 and 12.

20 75. A method according to claim 73, wherein said DME-related biallelic marker is selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284.

25

76. A method according to claim 73, wherein said haplotype comprises one of the following sets of biallelic markers:

12-455-326, 12-453-429 and 12-424-198;

12-455-326, 12-453-429, 12-424-198 and 12-454-363;

30 12-447-58, 12-455-326, 12-461-299 and 12-453-429;

12-441-233, 12-461-299 and 12-453-429;

12-441-233, 12-461-299, 12-453-429 and 12-426-154;

12-426-154, and 12-424-198;

12-426-154, 12-461-299, and 12-424-198;

**RECTIFIED SHEET (RULE 91)**

**ISA/EP**

158

- 10-428-219, and 12-724-225;  
12-128-225, 12-156-91, 12-139-380, and 12-140-134;  
12-148-311, 12-156-91, 12-139-380, 12-140-134;  
10-470-25, and 12-652-203; and  
5 10-470-25, 12-637-219, and 12-652-203.

77. A method according to claim 73, wherein said haplotype comprises one of the following sets of biallelic markers:

- 12-455-326, 12-453-429 and 12-424-198;  
10 12-455-326, 12-453-429, 12-424-198 and 12-454-363;  
12-447-58, 12-455-326, 12-461-299 and 12-453-429;  
12-426-154, and 12-424-198;  
10-428-219, and 12-724-225;  
12-128-225, 12-156-91, 12-139-380, and 12-140-134; and  
15 10-470-25, and 12-652-203.

78. A diagnostic kit comprising a polynucleotide according to any one of claims 2, 3, 4, 5, 10, 11, 13, 14, 15, and 16.

20 79. A polynucleotide for use in a hybridization assay for determining the identity of a nucleotide at a DME-related biallelic marker or MGST-II-related biallelic marker.

80. A polynucleotide for use in a sequencing assay for determining the identity of a nucleotide at a DME-related biallelic marker or MGST-II-related biallelic marker.

25

81. A polynucleotide for use in an allele specific amplification assay for determining the identity of a DME-related biallelic marker or MGST-II-related biallelic marker.

82. A polynucleotide for use in amplifying a segment of nucleotides comprising a DME-  
30 related biallelic marker or MGST-II-related biallelic marker.

83. A use according to any one of claims 79 to 82. wherein said polynucleotide is selected from the sequences described in Figure 1.

84. A method according to claim 83, wherein said DME-related biallelic marker is selected from the group consisting of the biallelic markers found in Figures 9, 10, 11 and 12.

5 85. A method according to claim 83, wherein said DME-related biallelic marker is selected from the group consisting of 12-455-326, 12-453-429, 12-454-363, 12-441-233, 12-461-299, 12-426-154, 12-424-198, 12-716-295, 10-428-219, 12-720-80, 12-156-91, 12-140-134, 12-653-423, 10-471-84, 10-471-85, 10-470-25, 12-652-203, 12-637-219, 12-721-440, and 10-420-284.

10

86. A method according to claim 83, wherein said haplotype comprises one of the following sets of biallelic markers:

12-455-326, 12-453-429 and 12-424-198;

12-455-326, 12-453-429, 12-424-198 and 12-454-363;

15

12-447-58, 12-455-326, 12-461-299 and 12-453-429;

12-441-233, 12-461-299 and 12-453-429;

12-441-233, 12-461-299, 12-453-429 and 12-426-154;

12-426-154, and 12-424-198;

12-426-154, 12-461-299, and 12-424-198;

20

10-428-219, and 12-724-225;

12-128-225, 12-156-91, 12-139-380, and 12-140-134;

12-148-311, 12-156-91, 12-139-380, 12-140-134;

10-470-25, and 12-652-203; and

10-470-25, 12-637-219, and 12-652-203.

25

87. A method according to claim 83, wherein said haplotype comprises one of the following sets of biallelic markers:

12-455-326, 12-453-429 and 12-424-198;

12-455-326, 12-453-429, 12-424-198 and 12-454-363;

30

12-447-58, 12-455-326, 12-461-299 and 12-453-429;

12-426-154, and 12-424-198;

10-428-219, and 12-724-225;

12-128-225, 12-156-91, 12-139-380, and 12-140-134; and

10-470-25, and 12-652-203.

160

88. An isolated, purified or recombinant, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of the following nucleotide positions of SEQ ID No. 485: 1-7466, 7726-20255, 20356-36904,  
5 36976-45166, 45249-45727 and 45966-49312.
89. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at  
10 position 36985, a C at position 45228 or a T at position 45755 of SEQ ID No. 485.
90. An isolated, purified, or recombinant polynucleotides comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID  
15 No. 486.
91. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at  
20 position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486.
92. An isolated, purified, or recombinant polynucleotides comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID  
25 No. 487.
93. An isolated, purified or recombinant polynucleotide comprising a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at  
30 position 326, a C at position 378 or a T at position 426 of SEQ ID No. 487.
94. An isolated, purified, or recombinant polynucleotides comprising a contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of SEQ ID Nos. 1-30, 436-441, 469-472, 474-477 and 484.

95. An isolated, purified, or recombinant polynucleotide comprising a contiguous span of 8 to 50 nucleotides of any one of SEQ ID Nos. 1-30, 436-441, 469-472, 474-477 and 484-487, or the complements thereof, wherein said contiguous span comprises a MGST-II-related biallelic marker.

96. A polynucleotide according to claim 95, wherein said MGST-II-related biallelic marker is selected from the group consisting of the biallelic markers described in Table 1; and the complements thereof.

10

97. A polynucleotide according to claim 95, wherein said MGST-II-related biallelic marker is selected from the group consisting of biallelic markers: 12-421-135, 12-421-140, 12-430-80, 12-441-233, 12-442-133, 12-447-58, 12-455-326, 12-461-299, 12-453-429, 12-424-198, 12-454-363, 12-458-196 and 12-426-154; and the complements thereof.

15

98. An isolated, purified, or recombinant polynucleotide which encodes a polypeptide comprising a contiguous span of at least 6 amino acids of SEQ ID No. 488, wherein said contiguous span includes a histidine residue at amino acid position 93 in SEQ ID No. 488.

99. An isolated, purified, or recombinant polynucleotide which encodes a polypeptide consisting essentially of amino acid residues 1-108 of SEQ ID No. 488.

100. An isolated, purified, or recombinant polynucleotide which encodes a polypeptide comprising a contiguous span of at least 6 amino acids of SEQ ID No. 489, wherein said contiguous span includes at least one of amino acid positions 20-30 of SEQ ID No. 489.

101. A recombinant vector comprising a polynucleotide according to any one of claims 88-95 and 98-100.

102. A host cell comprising a recombinant vector according to claim 101.

103. A non-human host animal or mammal comprising a recombinant vector according to claim 101.

104. A mammalian host cell comprising an MGST-II gene disrupted by homologous recombination with a knock out vector, comprising a polynucleotide according to any one of claims 88-95 and 98-100.
- 5 105. A non-human host mammal comprising a MGST-II gene disrupted by homologous recombination with a knock out vector, comprising a polynucleotide according to any one of claims 88-95 and 98-100.
106. An isolated, purified, or recombinant polypeptide comprising a contiguous span of at  
10 least 6 amino acids of SEQ ID No. 488, wherein said contiguous span includes a histidine residue at amino acid position 93 of SEQ ID No. 488.
107. An isolated or purified antibody composition capable of selectively binding to an epitope-containing fragment of a polypeptide according to claim 106, wherein said epitope  
15 comprises a histidine residue at amino acid position 93 in SEQ ID No. 488.
108. An isolated, purified, or recombinant polypeptide consisting essentially of amino acid residues 1-108 of SEQ ID No. 488.
- 20 109. An isolated, purified, or recombinant polypeptide comprising a contiguous span of at least 6 amino acids of SEQ ID No. 489, wherein said contiguous span includes at least one of amino acid positions 20-30 of SEQ ID No. 489.
110. An isolated or purified antibody composition capable of selectively binding to an  
25 epitope-containing fragment of a polypeptide according to claim 109, wherein said epitope comprises at least one of amino acid positions 20-30 of SEQ ID No. 489.
111. A method of determining whether an individual is at risk of developing asthma, or whether said individual suffers from asthma, comprising:
- 30 a) genotyping said individual for at least one MGST-II-related biallelic marker according to the method of claim 24; and
- b) correlating the result of step a) with a risk of developing asthma.

112. A method of determining whether an individual is at risk of developing hepatotoxicity upon treatment with zileuton, comprising:

a) genotyping said individual for at least one MGST-II-related biallelic marker according to the method of claim 24; and

5 b) correlating the result of step a) with a risk of developing asthma.

113. A method according to any one of claims 111 and 112, wherein said MGST-II-related biallelic marker is selected from the group consisting of biallelic markers: 12-455-326, 12-453-429, 12-424-198, 12-454-363, 12-447-58, 12-461-299, 12-441-233, and 12-10 426-154.

114. A diagnostic kit comprising a polynucleotide according to any one of claims 20, 23 and 88-97.

15 115. A computer readable medium having stored thereon a sequence selected from the group consisting of a nucleic acid code comprising one of the following:

a) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of the following nucleotide positions of SEQ ID No. 485: 1-7667, 7726-20264, 20365-36918, 20 36991-45180, 45263-45741 and 45980-49327;

b) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 36985, a C at position 45228 or a T at position 45755 of SEQ ID No. 485;

25 c) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 486;

d) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide 30 selected from the group consisting of a T at position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486;

e) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 487;

164

- f) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 325, a C at position 378 or a T at position 426 of SEQ ID No. 487;
- 5 g) contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of SEQ ID Nos. 2, 3, 5-30, 437-441, 472;
- h) a contiguous span of 8 to 50 nucleotides of any one of SEQ ID Nos 1 to 3 and 8 to 44, or the complements thereof, wherein said contiguous span comprises a MGST-II-related biallelic marker.
- 10 i) a nucleotide sequence complementary to any one of the contiguous spans of a), b), c), d), e), f), g) and h).

116. A computer readable medium having stored thereon a sequence consisting of a polypeptide code comprising one of the following:

- 15 a) a contiguous span of at least 6 amino acids of SEQ ID No. 488, wherein said contiguous span includes a histidine residue at amino acid position 93 of SEQ ID No. 488;
- b) a polypeptide consisting essentially of amino acid residues 1-108 of SEQ ID No. 488;
- 20 c) a contiguous span of at least 6 amino acids of SEQ ID No. 489, wherein said contiguous span includes at least one of amino acid positions 20-30 of SEQ ID No. 489.

117. A computer system comprising a processor and a data storage device wherein said  
25 data storage device a computer readable medium according to with claim 115 or 116.

118. A computer system according to claim 117, further comprising a sequence comparer and a data storage device having reference sequences stored thereon.

30 119. A computer system of claim 118 wherein said sequence comparer comprises a computer program which indicates polymorphisms.

120. A computer system of claim 118 further comprising an identifier which identifies features in said sequence.



121. A method for comparing a first sequence to a reference sequence, comprising the steps of:

a) reading said first sequence and said reference sequence through use of a computer  
5 program which compares sequences; and

b) determining differences between said first sequence and said reference sequence with said computer program,

wherein said first sequence is selected from the group consisting of a nucleic acid code comprising one of the following:

10 1) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of the following nucleotide positions of SEQ ID No. 485: 1-7667, 7726-20264, 20365-36918, 36991-45180, 45263-45741 and 45980-49327;

15 2) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 36985, a C at position 45228 or a T at position 45755 of SEQ ID No. 485;

20 3) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 486;

4) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486;

25 5) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 487;

30 6) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 325, a C at position 378 or a T at position 426 of SEQ ID No. 487;

7) contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of SEQ ID Nos. 2, 3, 5-30, 437-441, 472;

166

8) a contiguous span of 8 to 50 nucleotides of any one of SEQ ID Nos. 1 to 3 and 8 to 44, or the complements thereof, wherein said contiguous span comprises a MGST-II-related biallelic marker.

5 9) a nucleotide sequence complementary to any one of the contiguous spans of 1-8.

and a polypeptide code comprising one of the following:

10) a contiguous span of at least 6 amino acids of SEQ ID No. 488, wherein said contiguous span includes a histidine residue at amino acid position 93 of SEQ ID No. 488;

10 11) a polypeptide consisting essentially of amino acid residues 1-108 of SEQ ID No. 488;

12) a contiguous span of at least 6 amino acids of SEQ ID No. 489, wherein said contiguous span includes at least one of amino acid positions 20-30 of SEQ ID No. 489.

15

122. A method according to Claim 121, wherein said step of determining differences between the first sequence and the reference sequence comprises identifying at least one polymorphism.

20 123. A method for identifying a feature in a sequence, comprising the steps of:

a) reading said sequence through the use of a computer program which identifies features in sequences; and

b) identifying features in said sequence with said computer program;

wherein said sequence is selected from the group consisting of a nucleic acid code

25 comprising one of the following:

1) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of the following nucleotide positions of SEQ ID No. 485: 1-7667, 7726-20264, 20365-36918, 36991-45180, 45263-45741 and 45980-49327;

30

2) a contiguous span of at least 12 nucleotides of SEQ ID No. 485 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 36985, a C at position 45228 or a T at position 45755 of SEQ ID No. 485;

167

3) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 486;

5 4) a contiguous span of at least 12 nucleotides of SEQ ID No. 486 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 426, a C at position 478 or a T at position 526 of SEQ ID No. 486;

10 5) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises at least 1 of nucleotide positions 1-198 of SEQ ID No. 487;

6) a contiguous span of at least 12 nucleotides of SEQ ID No. 487 or the complementary sequence thereof, wherein said contiguous span comprises a nucleotide selected from the group consisting of a T at position 325, a C at position 378 or a T at position 426 of SEQ ID No. 487;

15 7) contiguous span of at least 12 nucleotides of a sequence selected from the group consisting of SEQ ID Nos 2, 3, 5-30, 437-441, 472;

8) a contiguous span of 8 to 50 nucleotides of any one of SEQ ID Nos. 1 to 3 and 8 to 44, or the complements thereof, wherein said contiguous span comprises a MGST-II-related biallelic marker.

20 9) a nucleotide sequence complementary to any one of the contiguous spans of 1-8.

and a polypeptide code comprising one of the following:

25 10) a contiguous span of at least 6 amino acids of SEQ ID No. 488, wherein said contiguous span includes a histidine residue at amino acid position 93 of SEQ ID No. 488;

11) a polypeptide consisting essentially of amino acid residues 1-108 of SEQ ID No. 488;

30 12) a contiguous span of at least 6 amino acids of SEQ ID No. 489, wherein said contiguous span includes at least one of amino acid positions 20-30 of SEQ ID No. 489.

Fig. 1

GENE	BIALLELIC MARKER ID	SEQ ID NO.	BIALLELIC MARKER POSITION IN SEQ ID NO.	VALIDATION MICRO- SEQUENCING	GENOTYPING LEAST COMMON ALLELE FREQUENCY %	
MGST2	12-421-140	1	501	N		
MGST2	12-424-192	2	501	N		
MGST2	12-424-198	3	501	N		
MGST2	12-425-57	4	501	N		
MGST2	12-426-154	5	461	N		
MGST2	12-429-198	6	501	N		
MGST2	12-430-80	7	501	Y	T	5.38
MGST2	12-433-215	8	501	N		
MGST2	12-441-233	9	501	Y	G	34.48
MGST2	12-441-343	10	501	N		
MGST2	12-442-221	11	501	N		
MGST2	12-447-58	12	501	Y	G	34.27
MGST2	12-453-429	13	501	Y	T	42.55
MGST2	12-454-363	14	501	N		
MGST2	12-455-326	15	501	Y	C	41.94
MGST2	12-455-383	16	501	N		
MGST2	12-456-269	17	501	N		
MGST2	12-456-380	18	501	N		
MGST2	12-457-204	19	501	N		
MGST2	12-457-206	20	501	N		
MGST2	12-458-196	21	501	N		
MGST2	12-458-438	22	501	N		
MGST2	12-460-274	23	501	N		
MGST2	12-461-124	24	501	N		
MGST2	12-461-299	25	501	Y	C	42.39
MGST2	12-461-465	26	501	N		
MGST2	12-462-280	27	501	N		
MGST2	12-464-66	28	501	N		
MGST2	12-465-26	29	501	N		
MGST2	12-465-234	30	501	N		
ME1	10-428-219	31	501	N		
ME1	10-429-84	32	501	N		
ME1	10-420-284	33	501	N		
ME1	10-423-411	34	501	N		
ME1	12-713-95	35	232	Y	C	31.18
ME1	12-713-149	36	286	N		
ME1	12-716-295	37	500	Y	T	32.8
ME1	12-720-80	38	480	Y	C	33.69
ME1	12-721-281	40	426	N		
ME1	12-721-440	41	501	Y	A	1.6
ME1	12-723-293	42	503	Y	T	0.54
ME1	12-724-195	43	501	N		
ME1	12-724-225	44	501	Y	T	25.86
CYP1A2	10-153-329	45	330	Y	G	2.69

Fig. 1 (following)

CYP1A2	10-95-342	46	342	Y	A	0.54
CYP1A2	10-100-277	47	277	Y	C	2.33
CYP1A2	10-102-294	48	297	Y	C	30.85
CYP2C8	10-413-394	49	501	Y	A	38.07
CYP2C8	10-414-243	50	501	N		
CYP2C8	10-416-273	51	501	Y	T	39.25
CYP2C8	10-418-177	52	501	Y	G	11.41
CYP2C8	12-665-315	53	501	Y	T	22.58
CYP2C8	12-666-324	54	501	Y	C	10.87
CYP2C9	10-76-177	56	178	N		
CYP2C9	10-76-217	57	218	N		
CYP2C9	10-76-333	58	334	Y	G	13.04
CYP2C9	10-77-316	59	501	N		
CYP2C9	10-155-78	60	78	N		
CYP2C9	10-155-104	61	104	N		
CYP2C9	10-156-52	62	52	Y	A	4.79
CYP2C9	10-157-39	63	39	Y	C	30.65
CYP2C9	10-157-131	64	131	Y		
CYP2C9	10-157-166	65	166	N		
CYP2C9	10-157-246	66	246	N		
CYP2C9	10-159-161	67	161	Y	A	11.29
CYP2C9	10-159-162	68	162	N		
CYP2C18	10-83-169	69	169	Y	T	12.9
CYP2C18	10-84-152	70	152	N		
CYP2C18	10-84-243	71	243	Y	T	32.56
CYP2C18	10-84-277	72	277	N		
CYP2C18	10-84-295	73	295	Y		
CYP2C18	10-85-43	74	43	Y		
CYP2C18	10-85-117	75	117	Y	T	12.35
CYP2C18	10-85-320	76	320	N		
CYP2C18	10-86-121	77	121	Y		
CYP3A4- CYP3A7	12-244-275	78	501	N		
CYP3A4- CYP3A7	12-251-153	79	450	Y		
CYP3A4- CYP3A7	12-254-115	80	246	N		
CYP3A4- CYP3A7	12-254-180	81	311	Y		
CYP3A4- CYP3A7	12-265-300	82	499	N		
CYP3A4- CYP3A7	12-271-118	83	501	N		
CYP3A4- CYP3A7	12-272-112	84	501	Y		
CYP3A7	10-216-182	85	503	Y	A	6.59
CYP3A7	10-217-91	86	501	Y	T	12.64
CYP3A7	10-213-292	87	503	Y	G	12.9
CYP3A7	10-214-279	88	501	Y	C	13.3
CYP3A7	10-214-380	89	501	Y	G	13.44
FMO	2-1-216	90	216	N		

Fig. 1 (following)

FMO	2-1-397	91	397	N		
FMO	2-3-232	92	232	N		
FMO	2-4-51	93	51	N		
FMO	2-4-126	94	126	N		
FMO	2-5-202	95	202	N		
FMO	2-5-275	96	275	N		
FMO	2-5-346	97	346	N		
FMO	2-8-171	98	171	N		
FMO	2-9-188	99	188	N		
FMO	2-9-223	100	223	N		
FMO	2-10-107	101	107	N		
FMO	2-10-378	102	377	N		
FMO	2-11-284	103	284	N		
FMO	2-11-156	104	156	N		
FMO	2-11-379	105	379	N		
FMO	2-12-223	106	223	N		
FMO	2-14-239	107	239	N		
FMO	2-14-370	108	370	N		
FMO	2-17-104	109	104	N		
FMO	2-17-396	110	395	N		
FMO	2-22-43	111	43	N		
FMO	2-22-138	112	138	N		
FMO	2-23-82	113	82	N		
FMO	2-23-166	114	166	N		
FMO	2-23-244	115	244	N		
FMO	2-24-115	116	115	N		
FMO	2-25-36	117	36	N		
FMO	2-27-185	118	185	N		
FMO	2-27-378	119	378	N		
FMO	2-29-142	120	142	N		
FMO	2-29-166	121	166	N		
FMO	2-29-205	122	204	N		
FMO	2-29-206	123	205	N		
FMO	2-29-314	124	313	N		
FMO	2-32-68	125	68	N		
FMO	2-35-357	126	357	N		
FMO	2-36-256	127	256	N		
FMO	2-36-354	128	353	N		
FMO	2-42-236	129	236	N		
FMO	2-43-139	130	139	N		
FMO	2-44-215	131	213	N		
FMO	2-45-38	132	38	N		
FMO	2-45-183	133	183	N		
FMO	2-45-335	134	335	N		
FMO	2-45-394	135	394	N		
FMO	2-48-39	136	39	N		
FMO	2-48-72	137	72	N		
FMO	2-48-156	138	156	N		
FMO	2-48-285	139	285	N		
FMO	2-49-167	140	167	N		
GSHR	10-436-43	141	501	Y	C	19.44

Fig. 1 (following)

GSHR	10-436-376	142	501	Y	A	32.76
GSHR	10-431-51	143	501	Y	A	20.21
GSHR	10-432-93	144	477	Y		
GSHR	12-631-208	145	433	Y	G	17.39
GSHS	10-260-282	146	501	N		
GSHS	10-263-26	147	503	N		
GSHS	10-258-408	148	501	N		
GSHS	12-317-259	149	501	N		
GSHS	12-323-385	150	501	Y	T	47.85
GSHS	12-324-219	151	357	N		
GSHS	12-324-335	152	473	Y	C	35.16
GSHS	12-324-380	153	501	N		
GSHS	12-325-30	154	501	N		
GSHS	12-327-31	155	479	Y	G	47.87
GSHS	12-327-415	156	500	N		
GSHS	12-331-270	157	501	N		
GSHS	12-331-275	158	501	N		
GSHS	12-334-320	159	503	N		
GSHS	12-334-391	160	503	N		
GSHS	12-335-417	161	501	N		
GSHS	12-337-189	162	501	N		
GSHS	12-340-130	163	381	N		
GSHS	12-340-210	164	461	Y	T	46.81
GSHS	12-340-222	165	473	N		
GSHS	12-340-240	166	491	N		
GSHS	12-341-99	167	501	Y	A	17.2
GSHS	12-342-32	168	501	N		
GSHS	12-344-349	169	501	Y	G	43.75
GSHS	12-345-453	170	501	N		
GSHS	12-346-204	171	501	N		
GLCL	10-364-55	172	501	N		
GLCL	10-364-108	173	501	N		
GLCL	10-364-267	174	501	N		
GLCL	10-367-20	175	497	N		
GLCL	10-351-389	176	501	N		
GLCL	10-353-102	177	501	N		
GLCL	12-474-346	178	501	Y	A	25.28
GLCL	10-354-320	179	501	N		
GLCL	10-354-360	180	501	N		
GLCL	10-355-87	181	501	N		
GLCL	10-358-60	182	501	N		
GLCL	12-468-63	183	501	Y	C	46.24
GLCL	12-468-388	184	501	N		
GLCL	12-468-491	185	501	N		
GLCL	12-469-132	186	501	N		
GLCL	12-469-245	187	501	N		
GLCL	12-472-435	188	501	N		
GLCL	12-473-311	189	501	N		
GLCL	12-473-483	190	501	N		
GLCL	12-475-85	191	501	N		
GLCL	12-475-446	192	485	N		

Fig. 1 (following)

GLCL	12-477-100	193	501	N		
GLCL	12-477-331	194	501	N		
GLCL	12-477-332	195	501	N		
GLCL	12-477-44	196	501	N		
GLCL	12-478-223	197	501	N		
GLCL	12-478-320	198	501	N		
GLCL	12-479-289	199	501	N		
GLCL	12-482-237	200	501	N		
GLCL	12-482-285	201	501	N		
GLCL	12-482-482	202	501	N		
GLCL	12-483-322	203	499	N		
GLCL	12-484-46	204	501	N		
GLCL	12-490-312	205	501	N		
GLCL	12-491-295	206	501	N		
GLCL	12-493-417	207	501	N		
GLCL	12-494-373	208	501	N		
GLCL	12-495-166	209	501	N		
GLCL	12-495-272	210	501	N		
GLCL	12-495-424	211	501	N		
GLCL	12-500-220	212	501	N		
GLCL	12-501-155	213	501	N		
GLCL	12-503-52	214	501	N		
GLCL	12-503-62	215	501	N		
GLCL	12-504-54	216	499	N		
GLCL	12-504-96	217	501	N		
GLCL	12-504-428	218	501	Y	C	47.22
GLCL	12-507-53	219	474	Y	C	47.78
GLCL	12-507-92	220	501	N		
GLCL	12-507-159	221	500	N		
GLCL	12-507-177	222	501	N		
GLCL	12-508-29	223	501	N		
GLCL	12-509-42	224	501	N		
GLCL	12-509-126	225	501	N		
GLCL	12-510-59	226	501	N		
GLCL	12-511-74	227	501	Y	C	11.45
GGT5	10-325-311	228	501	N		
GGT5	10-327-120	229	501	N		
GGT5	10-331-179	230	501	N		
GGT5	10-331-357	231	501	N		
GGT5	10-334-263	232	501	N		
GGT5	10-321-226	233	501	N		
GGT5	12-183-98	234	501	N		
GGT5	12-185-78	235	501	N		
GGT5	12-186-154	236	501	Y	A	6.99
GGT5	12-186-397	237	501	Y	C	28.08
GGT5	12-187-65	238	501	Y	C	23.37
GGT5	12-187-66	239	501	N		
GGT5	12-189-348	240	501	Y	A	26.09
GGT5	12-192-63	241	501	N		
GGT5	12-192-64	242	502	N		
GGT5	12-192-268	243	501	N		



Fig. 1 (following)

GGT5	12-192-334	244	501	N		
GGT5	12-192-352	245	501	N		
GGT5	12-194-135	246	501	N		
GGT5	12-194-325	247	501	N		
GGT5	12-194-337	248	501	N		
GGT5	12-194-479	249	501	N		
DP	10-442-133	250	501	Y	C	7.06
DP	10-444-248	251	484	Y	G	42.78
DP	10-445-281	252	501	Y	T	13.3
DP	12-668-362	253	501	N		
DP	12-670-48	254	501	N		
DP	12-670-91	255	501	N		
DP	12-670-157	256	501	Y	T	47.83
DP	12-671-148	257	501	Y	C	41.49
DP	12-679-245	258	253	N		
DP	12-679-371	259	379	N		
DP	12-679-426	260	434	N		
DP	12-680-331	261	503	N		
G6PDH	10-151-154	262	154	Y	A	0.54
G6PDH	10-138-206	263	205	Y	T	8.51
G6PDH	10-138-352	264	351	Y	C	14.04
PGDH	12-586-414	265	501	Y	A	28.33
PGDH	12-587-379	266	499	N		
PGDH	12-588-103	267	501	Y	A	48.88
PGDH	12-589-152	268	501	N		
PGDH	12-592-118	269	501	Y	A	49.46
PGDH	12-593-174	270	501	Y	C	34.71
PGDH	12-596-124	271	501	Y	A	26.7
PGDH	12-602-196	272	501	Y	C	31.72
PGDH	12-602-350	273	501	N		
PGDH	12-603-191	274	501	Y	C	27.42
PGDH	12-783-73	275	501	N		
PGDH	12-783-421	276	501	N		
PGDH	12-785-200	277	501	N		
PGDH	12-785-393	278	501	N		
PGDH	12-787-103	279	501	N		
PGDH	12-790-396	280	501	N		
PGDH	12-791-211	281	501	N		
PGDH	12-792-233	282	501	N		
PGDH	12-793-383	283	505	N		
PGDH	12-803-125	284	501	N		
PGDH	12-805-115	285	501	N		
PGDH	12-808-52	286	501	N		
PGDH	12-808-75	287	501	N		
PGDH	12-809-119	288	501	N		
PGDH	12-810-77	289	501	N		
PGDH	10-265-178	290	501	N		
PGDH	10-266-203	291	503	N		
UGT1A7	10-403-312	292	501	N		
UGT1A7	10-405-54	293	501	N		
UGT1A7	10-408-356	294	501	N		

Fig. 1 (following)

UGT1A7	10-409-148	295	501	N		
UGT1A7	10-409-249	296	501	N		
UGT1A7	10-410-274	297	501	N		
UGT1A7	10-410-280	298	501	N		
UGT1A7	10-410-337	299	501	N		
UGT1A7	12-121-326	300	501	Y	A	26.88
UGT1A7	12-122-341	301	501	N		
UGT1A7	12-122-381	302	501	N		
UGT1A7	12-124-169	303	501	N		
UGT1A7	12-124-194	304	501	Y	C	19.41
UGT1A7	12-124-300	305	501	N		
UGT1A7	12-124-58	306	501	N		
UGT1A7	12-126-222	307	501	N		
UGT1A7	12-126-297	308	501	N		
UGT1A7	12-128-225	309	501	Y	G	47.44
UGT1A7	12-129-176	310	501	N		
UGT1A7	12-130-203	311	501	N		
UGT1A7	12-130-260	312	501	N		
UGT1A7	12-131-112	313	501	N		
UGT1A7	12-132-157	314	255	N		
UGT1A7	12-132-437	315	501	N		
UGT1A7	12-133-153	316	666	N		
UGT1A7	12-133-318	317	501	N		
UGT1A7	12-136-238	318	249	N		
UGT1A7	12-138-141	319	501	N		
UGT1A7	12-138-42	320	501	N		
UGT1A7	12-138-67	321	501	N		
UGT1A7	12-139-380	322	501	Y	A	21.24
UGT1A7	12-140-134	323	501	Y	T	41.01
UGT1A7	12-140-329	324	501	N		
UGT1A7	12-140-385	325	501	N		
UGT1A7	12-141-159	326	501	Y	T	37.5
UGT1A7	12-141-392	327	501	N		
UGT1A7	12-142-315	328	501	N		
UGT1A7	12-142-321	329	501	Y	G	45.56
UGT1A7	12-143-453	330	501	Y	G	49.39
UGT1A7	12-144-169	331	501	N		
UGT1A7	12-144-33	332	501	N		
UGT1A7	12-146-174	333	501	N		
UGT1A7	12-146-47	334	501	N		
UGT1A7	12-148-283	335	501	N		
UGT1A7	12-148-311	336	501	Y	T	43.17
UGT1A7	12-149-320	337	501	N		
UGT1A7	12-151-174	338	501	N		
UGT1A7	12-151-196	339	501	N		
UGT1A7	12-151-270	340	501	N		
UGT1A7	12-152-453	341	501	N		
UGT1A7	12-153-116	342	501	Y	T	37.78
UGT1A7	12-154-480	343	501	N		
UGT1A7	12-155-403	344	501	N		
UGT1A7	12-156-91	345	501	Y	A	50

Fig. 1 (following)

UGT1A7	12-157-437	346	501	N		
UGT1A7	12-158-213	347	501	N		
UGT1A7	12-158-450	348	480	N		
UGT1A7	12-161-157	349	501	N		
UGT1A7	12-162-21	350	498	N		
UGT2B4	10-470-25	351	503	Y	T	44.44
UGT2B4	10-471-84	352	503	Y	A	27.17
UGT2B4	10-471-85	353	503	Y		
UGT2B4	10-472-202	354	503	Y	C	9.68
UGT2B4	10-473-333	355	503	N		
UGT2B4	10-494-284	356	503	N		
UGT2B4	12-637-219	357	499	Y	G	36.17
UGT2B4	12-639-95	358	499	Y	G	34.95
UGT2B4	12-639-241	359	499	N		
UGT2B4	12-640-151	360	499	N		
UGT2B4	12-640-296	361	499	N		
UGT2B4	12-640-325	362	499	Y		
UGT2B4	12-640-413	363	499	N		
UGT2B4	12-641-120	364	499	N		
UGT2B4	12-641-122	365	499	N		
UGT2B4	12-641-223	366	432	N		
UGT2B4	12-641-267	367	388	N		
UGT2B4	12-642-387	368	503	N		
UGT2B4	12-642-417	369	503	Y	G	26.51
UGT2B4	12-646-429	370	434	N		
UGT2B4	12-646-433	371	438	N		
UGT2B4	12-647-145	372	503	N		
UGT2B4	12-648-123	373	251	N		
UGT2B4	12-648-300	374	428	Y	T	37.91
UGT2B4	12-648-402	375	501	N		
UGT2B4	12-652-115	376	298	N		
UGT2B4	12-652-203	377	386	Y	C	42.47
UGT2B4	12-652-274	378	457	N		
UGT2B4	12-652-371	379	501	N		
UGT2B4	12-653-423	380	499	N		
UGT2B4	12-654-115	381	499	N		
UGT2B4	12-654-207	382	499	N		
UGT2B4	12-657-396	383	503	N		
UGT2B4	12-658-120	384	503	N		
UGT2B4	12-659-382	385	501	N		
UGT2B4	12-660-134	386	306	N		
UGT2B4	12-662-80	387	497	N		
UGT2B7	12-906-149	388	501	Y		
UGT2B7	12-906-154	389	501	N		
UGT2B7	12-906-251	390	501	N		
UGT2B7	12-906-451	391	501	N		
UGT2B7	12-907-199	392	244	Y	G	3.23
UGT2B7	12-907-482	393	501	N		
UGT2B7	12-909-36	394	53	N		
UGT2B7	12-909-176	395	193	N		
UGT2B7	12-909-484	396	501	N		

Fig. 1 (following)

UGT2B7	12-910-76	397	347	Y	A	23.91
UGT2B7	12-910-295	398	503	N		
UGT2B7	12-911-22	399	240	Y	C	43.96
UGT2B7	12-912-65	400	501	N		
UGT2B7	12-914-106	401	384	Y	C	44.62
UGT2B7	12-914-252	402	503	N		
UGT2B10	10-448-266	403	503	N		
UGT2B10	10-453-330	404	503	N		
UGT2B10	10-455-367	405	503	N		
UGT2B10	12-5-158	406	503	Y	T	11.8
UGT2B10	12-9-367	407	499	Y	A	14.67
UGT2B10	12-10-303	408	501	Y	T	14.67
UGT2B10	12-14-264	409	501	Y	T	14.29
UGT2B10	12-17-86	410	499	Y	A	44.02
UGT2B10	12-19-163	411	501	Y	G	29.89
UGT2B15	10-457-284	412	503	N		
UGT2B15	10-460-221	413	503	N		
UGT2B15	10-460-232	414	503	N		
UGT2B15	10-460-235	415	503	N		
UGT2B15	10-460-236	416	503	N		
UGT2B15	10-460-285	417	503	N		
UGT2B15	12-605-58	418	501	Y	T	47.67
UGT2B15	12-607-207	419	501	N		
UGT2B15	12-609-119	420	499	Y	T	45.11
UGT2B15	12-609-180	421	499	N		
UGT2B15	12-609-233	422	499	N		
UGT2B15	12-611-294	423	501	Y	A	43.26
UGT2B15	12-612-41	424	501	Y	C	28.65
UGT2B15	12-613-302	425	499	N		
UGT2B15	12-614-471	426	501	Y	T	39.44
UGT2B15	12-620-192	427	503	Y	T	30.36
UGT2B15	12-621-49	428	503	N		
UGT2B15	12-622-325	429	364	N		
UGT2B15	12-624-82	430	501	N		
UGT2B15	12-624-83	431	501	N		
UGT2B15	12-624-107	432	489	N		
UGT2B15	12-624-146	433	501	N		
UGT2B15	12-624-288	434	501	N		
UGT2B15	12-624-293	435	501	N		
MGST2	12-421-135	436	501	N		
MGST2	12-442-133	437	501	Y	C	5.85
MGST2	12-449-63	438	501	N		
MGST2	12-454-242	439	501	N		
MGST2	12-463-250	440	501	N		
MGST2	12-462-199	441	501	N		
DME	10-430-287	442	501	N		
DME	12-718-432	443	501	N		
CYP3A4- CYP3A7	12-269-301	444	501	Y		
FMO	2-13-398	445	501	N		
FMO	2-28-132	446	501	N		

Fig. 1 (following)

FMO	2-39-27	447	501	N		
FMO	2-45-155	448	501	N		
FMO	2-4-391	14	501	N		
GSHS	12-345-410	450	501	N		
GLCL	10-358-353	451	501	N		
GLCL	10-360-190	452	501	N		
GLCL	10-365-374	453	501	N		
GLCL	10-367-58	454	501	N		
GLCL	12-468-424	455	501	N		
GLCL	12-481-293	456	501	N		
GLCL	12-499-86	457	501	N		
GLCL	12-500-217	458	501	N		
GLCL	12-511-101	459	501	N		
6PGD	12-586-443	460	501	N		
6PGD	12-593-287	461	501	N		
6PGD	12-795-383	462	501	N		
UGT2B4	10-494-332	463	501	N		
UGT2B4	12-659-251	465	429	N		
UGT2B7	12-912-419	466	501	N		
UGT2B7	12-914-28	467	306	N		
UGT2B15	12-624-307	468	501	N		
MGST2	10-290-326	469	501	N		
MGST2	10-290-37	470	501	N		
MGST2	10-523-232	471	501	N		
MGST2	12-449-300	472	501	N		
G6PDH	10-186-212	473	212	N		
MGST2	10-286-289	474	501	N		
MGST2	10-286-345	475	501	N		
MGST2	10-286-375	476	501	N		
MGST2	10-289-201	477	501	N		
GGT5	10-321-28	478	501	N		
CYP1A2	10-98-265	479	265	N		
UGT1A7	12-157-115	480	501	N		
GLCL	12-472-48	481	501	N		
GLCL	12-477-151	482	501	N		
GLCL	12-479-214	483	501	N		
MGST2	10-290-328	484	501	N		

11/65

**FIG. 2**

SEQ ID NO.	BIALLELIC MARKER ID	1 <sup>ST</sup> ALLELE	2 <sup>ND</sup> ALLELE	POSITION RANGE OF PREFERRED SEQUENCES
1	12-421-140	A	G	1-1001
2	12-424-192	A	G	190-801; 865-999
3	12-424-198	G	T	184-795; 859-993
4	12-425-57	G	A	208-225; 266-478
5	12-426-154	A	G	152-961
6	12-429-198	C	T	260-784; 822-1001
7	12-430-80	T	C	1-996
8	12-433-215	A	G	1-1001
12	12-447-58	G	C	1-36; 390-914
13	12-453-429	C	T	1-1001
14	12-454-363	A	G	1-315; 377-466; 598-619
15	12-455-326	T	C	1-357; 391-594; 760-827
16	12-455-383	G	A	1-414; 448-651; 817-884
17	12-456-269	A	G	1-536
19	12-457-204	A	G	437-527; 761-1001
20	12-457-206	C	T	435-525; 759-1001
21	12-458-196	T	A	1-727
22	12-458-438	T	C	1-21; 298-1001
23	12-460-274	A	G	1-499; 563-1001
24	12-461-124	A	C	1-203; 259-644; 687-773; 807-833
25	12-461-299	C	T	1-28; 84-469; 512-591
26	12-461-465	C	T	1-303; 346-432
27	12-462-280	C	T	1-1001
28	12-464-66	G	T	1-215; 261-1001
29	12-465-26	C	T	1-61; 99-1001
30	12-465-234	G	T	1-1001
31	10-428-219	A	G	1-398; 506-1000
33	10-420-284	C	T	1-302; 472-772
34	10-423-411	C	T	1-227; 333-821
35	12-713-95	C	T	1-668
36	12-713-149	G	C	1-668
37	12-716-295	C	T	1-902
38	12-720-80	G	C	1-982
40	12-721-281	A	C	1-926
41	12-721-440	A	G	1-1000
42	12-723-293	C	T	1-1001
43	12-724-195	C	T	277-1001
44	12-724-225	C	T	245-1001
49	10-413-394	A	G	1-139; 831-1001
51	10-416-273	A	T	1-258; 855-1001
53	12-665-315	T	C	180-1001
54	12-666-324	A	C	338-375; 414-501; 581-611

12/65

FIG. 2 (following)

78	12-244-275	A	G	
79	12-251-153	A	C	1-111; 158-950
82	12-265-300	T	C	1-999
84	12-272-112	A	C	396-416
85	10-216-182	A	G	77-151; 186-235; 320-488; 733-761
86	10-217-91	C	T	1-189; 434-462; 810-871; 925-1001
101	2-10-107	C	T	260-450
102	2-10-378	A	G	260-450
104	2-11-156	A	C	369-387
105	2-11-379	A	G	369-387
121	2-29-166	C	T	
141	10-436-43	G	C	1-630; 794-1001
142	10-436-376	A	G	1-297; 461-642; 977-1001
143	10-431-51	A	C	329-534; 682-860
144	10-432-93	A	G	1-40; 374-559; 705-885
146	10-260-282	G	T	1-24; 95-324; 482-1001
147	10-263-26	A	C	1-632; 754-1001
148	10-258-408	A	G	337-668
149	12-317-259	G	A	1-44; 116-150; 181-241; 281-673; 968-1001
150	12-323-385	T	C	340-1001
152	12-324-335	G	C	1-176; 395-906
153	12-324-380	A	G	1-159; 378-889
154	12-325-30	C	T	272-837; 977-1001
155	12-327-31	G	T	1-570; 655-821; 875-922; 973-999
156	12-327-415	A	G	1-208; 293-459; 513-1001
157	12-331-270	G	A	1-531; 834-1001
158	12-331-275	T	G	125-536; 839-1001
159	12-334-320	A	G	68-705
160	12-334-391	A	G	1-634; 982-1001
161	12-335-417	G	C	239-1001
162	12-337-189	A	G	1-1001
163	12-340-130	A	G	196-881
164	12-340-210	C	T	196-961
165	12-340-222	G	T	196-973
166	12-340-240	C	T	196-975
167	12-341-99	A	G	1-1001
168	12-342-32	T	C	238-397; 859-960
169	12-344-349	G	T	365-1001
170	12-345-453	G	C	1-606
192	12-475-446	G	A	
228	10-325-311	A	G	41-312; 421-891
231	10-331-357	G	T	76-218; 373-730; 938-1001
232	10-334-263	A	G	1-359; 481-604; 781-893
235	12-185-78	C	T	1-1001
236	12-186-154	A	G	335-876
237	12-186-397	C	T	92-633; 967-1001

13/65

FIG. 2 (following)

238	12-187-65	C	T	345-846
239	12-187-66	A	G	344-845
240	12-189-348	G	A	281-358; 407-1001
246	12-194-135	G	T	543-1001
247	12-194-325	A	G	351-1001
248	12-194-337	A	G	339-1001
249	12-194-479	C	T	197-1001
251	10-444-248	A	G	
254	12-670-48	G	C	1-961
255	12-670-91	C	T	1-918
256	12-670-157	C	T	1-852
257	12-671-148	C	T	1-908
258	12-679-245	A	G	96-465
259	12-679-371	A	G	96-465
260	12-679-426	C	T	96-465
265	12-586-414	A	G	1-71; 149-929
268	12-589-152	T	G	164-1001
269	12-592-118	A	T	353-1001
270	12-593-174	T	C	342-815
271	12-596-124	A	G	1-742
272	12-602-196	C	T	1-240; 436-641
274	12-603-191	T	C	1-709
275	12-783-73	G	C	1-769; 981-1001
277	12-785-200	C	T	351-510
279	12-787-103	G	A	1-47; 232-324; 401-1001
280	12-790-396	G	A	47-64; 393-1001
281	12-791-211	A	G	1-379; 467-818
284	12-803-125	T	A	125-1001
285	12-805-115	G	A	1-66; 278-838; 959-1001
286	12-808-52	A	G	400-1001
287	12-808-75	G	C	377-1001
289	12-810-77	G	A	99-1001
293	10-405-54	C	T	1-492
297	10-410-274	A	C	643-805
301	12-122-341	C	T	1-23; 150-282; 324-435; 593-620
302	12-122-381	A	C	110-242; 284-395; 553-580
303	12-124-169	G	T	1-727; 788-1001
304	12-124-194	C	T	1-702; 763-1001
305	12-124-300	G	T	1-596; 657-1001
306	12-124-58	A	C	1-837; 898-1001
308	12-126-297	T	C	1-508; 944-1001
310	12-129-176	G	T	163-1001
313	12-131-112	T	C	254-422
315	12-132-437	A	C	258-991
316	12-133-153	T	C	1-515; 607-666; 775-918; 976-1001
317	12-133-318	T	A	1-515; 607-666; 775-918; 976-1001



14/65

FIG. 2 (following)

318	12-136-238	A	G	1-643
319	12-138-141	G	A	1-393; 521-651
320	12-138-42	T	G	1-294; 422-552
321	12-138-67	G	A	1-319; 447-577
322	12-139-380	A	G	94-157; 276-663
323	12-140-134	G	T	1-142; 212-951
324	12-140-329	C	T	1-756
325	12-140-385	G	C	1-700
326	12-141-159	T	C	1-1001
327	12-141-392	C	A	1-1001
328	12-142-315	A	G	1-1001
329	12-142-321	A	G	1-1001
330	12-143-453	A	G	1-1001
331	12-144-169	G	A	97-1001
332	12-144-33	G	A	1-1001
333	12-146-174	T	C	1-529; 765-805
336	12-148-311	T	C	1-251; 684-1001
337	12-149-320	T	C	202-1001
338	12-151-174	G	T	313-1001
339	12-151-196	C	T	291-1001
340	12-151-270	A	G	217-1001
341	12-152-453	C	T	1-239; 526-576; 623-1001
342	12-153-116	C	T	1-819; 866-1001
346	12-157-437	A	C	1-865
347	12-158-213	T	C	1-1001
348	12-158-450	T	G	206-980
350	12-162-21	A	G	1-998
351	10-470-25	A	T	1-339; 413-539; 730-1001
355	10-473-333	C	T	461-744
356	10-494-284	C	T	89-349; 450-513
359	12-639-241	T	G	1-199; 652-667; 742-763
360	12-640-151	G	A	1-806; 943-1001
361	12-640-296	T	G	1-862
362	12-640-325	T	C	1-1001
363	12-640-413	C	G	1-1001
364	12-641-120	G	A	98-1001
365	12-641-122	T	G	100-1001
366	12-641-223	G	A	1-15; 134-1001
367	12-641-267	T	C	1-15; 134-1001
368	12-642-387	A	G	1-224; 292-1001
369	12-642-417	A	G	1-194; 262-1001
372	12-647-145	A	G	1-898
373	12-648-123	A	G	1-751
374	12-648-300	C	T	1-839
375	12-648-402	G	C	1-810; 909-1001
376	12-652-115	C	T	1-30; 191-798

15/65

FIG. 2 (following)

377	12-652-203	A	C	1-30; 191-886
378	12-652-274	G	T	1-30; 191-957
379	12-652-371	C	T	138-1001
380	12-653-423	T	A	1-603
381	12-654-115	T	C	161-1001
382	12-654-207	G	A	1-95; 253-1001
383	12-657-396	A	G	329-1001
384	12-658-120	A	T	1-1001
385	12-659-382	A	G	1-1001
386	12-660-134	A	G	291-608; 886-983
387	12-662-80	G	C	480-1001
391	12-906-451	A	C	447-532; 865-1001
392	12-907-199	G	T	
394	12-909-36	A	G	
397	12-910-76	A	G	1-785; 854-870
398	12-910-295	C	T	1-1001
399	12-911-22	G	C	1-740
400	12-912-65	C	T	1-183; 290-1001
401	12-914-106	C	T	1-1001
402	12-914-252	A	T	1-1001
404	10-453-330	C	T	1-207; 761-1001
406	12-5-158	C	T	672-1001
407	12-9-367	G	A	1-15; 249-283; 400-425; 493-666; 722-1001
409	12-14-264	C	T	1-1001
410	12-17-86	T	A	1-1001
411	12-19-163	A	G	230-388; 525-709; 824-866
414	10-460-232	A	G	1-349; 521-1001
415	10-460-235	C	T	1-346; 518-1001
416	10-460-236	A	G	1-345; 517-1001
417	10-460-285	A	T	1-296; 468-1001
418	12-605-58	G	T	1-1001
419	12-607-207	G	A	1-746; 885-902
420	12-609-119	T	G	1-234; 294-1001
421	12-609-180	T	G	40-295; 355-1001
422	12-609-233	G	A	138-348; 404-1001
423	12-611-294	G	A	1-617; 895-967
424	12-612-41	C	T	1-56; 398-708
425	12-613-302	C	G	1-86; 336-1001
426	12-614-471	T	A	1-1001
427	12-620-192	G	T	318-1001
428	12-621-49	A	G	1-242; 451-815
429	12-622-325	C	T	
430	12-624-82	T	C	1-1001
431	12-624-83	G	A	1-1001
432	12-624-107	T	C	1-989
433	12-624-146	T	C	1-1001

16/65

FIG. 2 (following)

434	12-624-288	T	G	1-1001
435	12-624-293	T	C	1-1001
436	12-421-135	T	-	1-1001
438	12-449-63	AT	-	1-107; 314-656
439	12-454-242	AT	-	1-436; 498-587; 719-740
440	12-463-250	CAT	-	1-30; 102-601; 752-1001
441	12-462-199		deletion	1-1001
442	10-430-287	T	-	1-330; 442-740; 782-1001
443	12-718-432	T	-	1-1001
444	12-269-301	T	-	774-820
450	12-345-410		deletion	1-649
460	12-586-443	C	-	1-42; 120-911
461	12-593-287		deletion	1-17; 455-942
462	12-795-383		insertion	166-500; 540-1001
465	12-659-251	deletion	-	1-929
466	12-912-419	A	-	1-686
467	12-914-28	T	-	1-806
468	12-624-307	T	-	1-1001
469	10-290-326	A	G	1-197; 437-1000
472	12-449-300	T	C	1-542; 908-1000
484	10-290-328	deletion		1-194; 434-1000

17/65

**Fig. 3**

SEQ ID NO.	BIALLELIC MARKER ID	ORIGINAL ALLELE	ALTERNATIVE ALLELE
10	12-441-343	G	A
18	12-456-380	G	T
32	10-429-84	T	C
45	10-153-329	T	G
46	10-95-342	G	A
47	10-100-277	T	C
57	10-76-217	C	T
59	10-77-316	A	T
60	10-155-78	T	C
61	10-155-104	C	G
62	10-156-52	G	A
63	10-157-39	C	T
64	10-157-131	A	G
65	10-157-166	G	A
66	10-157-246	G	A
67	10-159-161	T	A
68	10-159-162	C	A
69	10-83-169	T	C
70	10-84-152	C	T
71	10-84-243	C	T
74	10-85-43	C	T
76	10-85-320	A	T
81	12-254-180	G	A
83	12-271-118	T	C
87	10-213-292	C	G
90	2-1-216	G	A
91	2-1-397	T	C
92	2-3-232	C	T
93	2-4-51	A	C
94	2-4-126	A	G
95	2-5-202	A	G
96	2-5-275	G	A
97	2-5-346	C	T
98	2-8-171	A	G
99	2-9-188	G	A
100	2-9-223	T	G
103	2-11-284	A	G
106	2-12-223	A	T
107	2-14-239	C	T
108	2-14-370	C	G
109	2-17-104	A	G
110	2-17-396	A	C
111	2-22-43	A	G
112	2-22-138	G	A
113	2-23-82	A	G
114	2-23-166	G	A
115	2-23-244	G	T

18/65

FIG. 3 (following)

116	2-24-115	C	T
117	2-25-36	G	C
118	2-27-185	G	A
119	2-27-378	G	C
120	2-29-142	G	A
124	2-29-314	C	T
125	2-32-68	A	G
126	2-35-357	T	C
127	2-36-256	G	C
128	2-36-354	A	C
129	2-42-236	C	T
130	2-43-139	G	A
131	2-44-215	C	T
132	2-45-38	C	T
133	2-45-183	T	C
134	2-45-335	T	A
135	2-45-394	C	T
136	2-48-39	A	G
137	2-48-72	C	T
138	2-48-156	T	G
139	2-48-285	A	G
140	2-49-167	A	G
151	12-324-219	C	T
171	12-346-204	G	A
172	10-364-55	T	G
173	10-364-108	T	C
174	10-364-267	T	A
175	10-367-20	T	G
176	10-351-389	A	G
177	10-353-102	T	A
178	12-474-346	G	A
179	10-354-320	G	A
180	10-354-360	A	G
181	10-355-87	G	A
182	10-358-60	G	A
183	12-468-63	T	C
184	12-468-388	T	C
185	12-468-491	G	A
186	12-469-132	T	C
187	12-469-245	G	A
188	12-472-435	G	A
189	12-473-311	A	C
190	12-473-483	T	C
191	12-475-85	G	T
193	12-477-100	A	G
194	12-477-331	G	A
195	12-477-332	T	C
196	12-477-44	C	G
197	12-478-223	G	A
198	12-478-320	G	A
199	12-479-289	G	T

19/65

FIG. 3 (following)

200	12-482-237	A	G
201	12-482-285	T	A
202	12-482-482	A	T
203	12-483-322	A	T
204	12-484-46	G	A
205	12-490-312	A	T
206	12-491-295	A	G
207	12-493-417	G	C
208	12-494-373	C	T
209	12-495-166	C	T
210	12-495-272	A	T
211	12-495-424	T	C
212	12-500-220	A	G
213	12-501-155	G	T
214	12-503-52	T	G
215	12-503-62	T	A
216	12-504-54	G	A
217	12-504-96	C	T
218	12-504-428	G	C
219	12-507-53	T	C
220	12-507-92	A	G
221	12-507-159	T	G
222	12-507-177	C	G
223	12-508-29	G	A
224	12-509-42	G	A
225	12-509-126	G	A
226	12-510-59	G	A
227	12-511-74	T	C
229	10-327-120	C	T
230	10-331-179	G	A
233	10-321-226	A	G
234	12-183-98	G	A/T
241	12-192-63	C	G/T
243	12-192-268	G	C
245	12-192-352	G	A
250	10-442-133	G	C
252	10-445-281	C	T
253	12-668-362	T	A
261	12-680-331	C	T
262	10-151-154	G	A
264	10-138-352	T	C
266	12-587-379	A	C
267	12-588-103	G	A
273	12-602-350	C	A
278	12-785-393	G	A
283	12-793-383	T	G
288	12-809-119	C	G
290	10-265-178	G	A
292	10-403-312	C	T
298	10-410-280	C	T
299	10-410-337	A	G

20/65

FIG. 3 (following)

300	12-121-326	G	A
307	12-126-222	C	T
309	12-128-225	G	T
312	12-130-260	C	T
314	12-132-157	T	C
334	12-146-47	A	G
335	12-148-283	G	A
343	12-154-480	C	T
344	12-155-403	C	A
349	12-161-157	G	A
354	10-472-202	T	C
358	12-639-95	G	A
370	12-646-429	T	C
371	12-646-433	T	G
388	12-906-149	G	A
389	12-906-154	A	C
390	12-906-251	A	T
395	12-909-176	C	T
396	12-909-484	G	T
403	10-448-266	A	C
405	10-455-367	T	C
408	12-10-303	C	T
413	10-460-221	C	T
437	12-442-133	C	-
445	2-13-398	G	-
446	2-28-132	-	T
447	2-39-27		deletion
448	2-45-155		deletion
449	2-4-391	G	-
452	10-360-190	-	T
453	10-365-374	-	A
454	10-367-58	-	insertion
455	12-468-424	-	T
457	12-499-86		deletion
458	12-500-217		insertion
459	12-511-101	A	-
463	10-494-332	insertion	
470	10-290-37	C	T
471	10-523-232	C	T
473	10-186-212	G	C
474	10-286-289	C	G
475	10-286-345	A	T
476	10-286-375	A	G
477	10-289-201	T	C
478	10-321-28	G	A
479	10-98-265	A	G
480	12-157-115	T	C
481	12-472-48	T	G
482	12-477-151	A	G
483	12-479-214	G	T

**Fig. 4**

SEQ ID NO.	BIALLELIC MARKER ID	1 <sup>ST</sup> ALLELE	2 <sup>ND</sup> ALLELE
9	12-441-233	G	A
11	12-442-221	T	C
48	10-102-294	C	T
50	10-414-243	A	G
52	10-418-177	A	G
56	10-76-177	A	T
58	10-76-333	G	C
72	10-84-277	A	G
73	10-84-295	A	G
75	10-85-117	G	T
77	10-86-121	A	C
80	12-254-115	A	T
88	10-214-279	C	T
89	10-214-380	A	G
122	2-29-205	C	T
123	2-29-206	A	G
145	12-631-208	A	G
242	12-192-64	T	C
244	12-192-334	G	A
263	10-138-206	C	T
276	12-783-421	C	T
282	12-792-233	G	A
291	10-266-203	C	T
294	10-408-356	C	T
295	10-409-148	G	C
296	10-409-249	G	C
311	12-130-203	C	T
345	12-156-91	A	G
352	10-471-84	A	T
353	10-471-85	A	C
357	12-637-219	G	A
393	12-907-482	A	T
412	10-457-284	G	T
451	10-358-353		deletion
456	12-481-293	T	-



22/65

**Fig. 5**

SEQ ID NO.	POSITION RANGE OF PREFERRED SEQUENCES
9	940-1001
18	1-425; 920-1001
32	1-175; 713-787
50	683-1001
52	1-450; 606-1001
58	732-832
59	612-725
87	717-742; 881-899; 951-1001
88	1-24; 276-305
103	369-387
129	433-449
145	152-241; 540-933
151	1-176; 395-857
171	1-98; 753-798
212	986-1001
229	1-439; 639-1001
230	254-396; 551-908
233	1-161; 283-406; 583-695; 843-1001
261	1-452
267	552-1001
273	1-86; 282-487; 963-1001
276	1-421; 633-1001
278	158-317; 834-1001
283	1-112
288	1-174; 252-399; 583-987
290	1-342; 790-1001
291	86-460
292	654-1001
298	637-799
299	580-742
300	1-297; 727-803
307	1-433; 869-1001
309	1-17
311	928-953
312	871-896; 972-1001
314	289-755
334	1-402; 638-678; 969-1001
335	1-223; 656-923
343	629-1000
349	98-251; 960-1001
352	1-52; 399-436
353	1-52; 399-436
354	754-1001
357	527-810
358	1-53; 506-521; 596-617
370	1-80; 675-1001
371	1-80; 675-1001

23/65

**FIG. 5 (following)**

388	1-293; 748-833
389	1-288; 743-828
390	1-191; 646-731
403	745-1001
405	984-1001
408	797-897; 931-1001
412	1-207; 979-1001
455	1-22
463	1-299; 400-463
470	1-485; 725-1000
471	1-432; 514-1000
474	1-353; 656-1000
475	1-298; 601-1000
476	1-268; 571-1000
477	1-452; 538-1000
479	385-391
480	910-954
482	443-531; 730-854
483	347-392

24/65

**Fig. 6**

SEQ ID NO.	POSITION RANGE OF MICROSEQUENCING PRIMERS	COMPLEMENTARY POSITION RANGE OF MICROSEQUENCING PRIMERS
1	481-500	502-521
2	481-500	502-521
3	481-500	502-521
4	481-500	502-521
5	441-460	462-481
6	481-500	502-521
7	482-500*	502-521
8	481-500	502-521
9	482-500*	502-521
10	481-500	502-521
11	481-500	502-521
12	482-500*	502-521
13	481-500	502-520*
14	481-500	502-521
15	481-500	502-520*
16	481-500	502-521
17	481-500	502-521
18	481-500	502-521
19	481-500	502-521
20	481-500	502-521
21	481-500	502-521
22	481-500	502-521
23	481-500	502-521
24	481-500	502-521
25	481-500	502-520*
26	481-500	502-521
27	481-500	502-521
28	481-500	502-521
29	481-500	502-521
30	481-500	502-521
31	481-500	502-521
32	481-500	502-521
33	481-500	502-521
34	481-500	502-521
35	213-231*	233-252
36	266-285	287-306
37	477-499*	501-520
38	461-479*	481-500
40	406-425	427-446
41	481-500	502-520*
42	484-502*	504-523
43	481-500	502-521
44	482-500*	502-521
45	310-329	331-349*
46	322-341	343-361*
47	258-276*	278-297
48	278-296*	298-317

25/65

FIG. 6 (following)

49	482-500*	502-520*
50	481-500	502-521
51	482-500*	502-520*
52	481-500	502-520*
53	482-500*	502-521
54	482-500*	502-521
56	158-177	179-198
57	198-217	219-238
58	315-333*	335-354
59	481-500	502-521
60	58-77	79-98
61	84-103	105-124
62	32-51	53-71*
63	20-38*	40-59
64	111-130	132-150*
65	146-165	167-186
66	226-245	247-266
67	142-160*	162-181
68	142-161	163-182
69	150-168*	170-189
70	132-151	153-172
71	224-242*	244-263
72	257-276	278-297
73	275-294	296-314*
74	24-42*	44-63
75	97-116	118-136*
76	300-319	321-340
77	102-120*	122-141
78	481-500	502-521
79	431-449*	451-470
80	226-245	247-266
81	292-310*	312-331
82	479-498	500-519
83	481-500	502-521
84	482-500*	502-521
85	483-502	504-522*
86	482-500*	502-521
87	484-502*	504-523
88	481-500	502-521
89	481-500	502-520*
90	196-215	217-236
91	377-396	398-417
92	212-231	233-252
93	31-50	52-71
94	106-125	127-146
95	182-201	203-222
96	255-274	276-295
97	326-345	347-366
98	151-170	172-191
99	168-187	189-208
100	203-222	224-243

26/65

FIG. 6 (following)

101	87-106	108-127
102	357-376	378-397
103	264-283	285-304
104	136-155	157-176
105	359-378	380-399
106	203-222	224-243
107	219-238	240-259
108	350-369	371-390
109	84-103	105-124
110	375-394	396-415
111	23-42	44-63
112	118-137	139-158
113	62-81	83-102
114	146-165	167-186
115	224-243	245-264
116	95-114	116-135
117	16-35	37-56
118	165-184	186-205
119	358-377	379-398
120	122-141	143-162
121	146-165	167-186
122	184-203	205-224
123	185-204	206-225
124	293-312	314-333
125	48-67	69-88
126	337-356	358-377
127	236-255	257-276
128	333-352	354-373
129	216-235	237-256
130	119-138	140-159
131	193-212	214-233
132	18-37	39-58
133	163-182	184-203
134	315-334	336-355
135	374-393	395-414
136	19-38	40-59
137	52-71	73-92
138	136-155	157-176
139	265-284	286-305
140	147-166	168-187
141	482-500*	502-521
142	481-500	502-520*
143	482-500*	502-521
144	457-476	481-499*
145	414-432*	434-453
146	481-500	502-521
147	483-502	504-523
148	481-500	502-521
149	481-500	502-521
150	481-500	502-520*
151	337-356	358-377

27/65

FIG. 6 (following)

152	454-472*	474-493
153	481-500	502-521
154	481-500	502-521
155	459-478	480-498*
156	480-499	501-520
157	481-500	502-521
158	481-500	502-521
159	483-502	504-523
160	483-502	504-523
161	481-500	502-521
162	481-500	502-521
163	361-380	382-401
164	442-460*	462-481
165	453-472	474-493
166	471-490	492-511
167	481-500	502-520*
168	481-500	502-521
169	481-500	502-520*
170	481-500	502-521
171	481-500	502-521
172	481-500	502-521
173	481-500	502-521
174	481-500	502-521
175	477-496	498-517
176	481-500	502-521
177	481-500	502-521
178	482-500*	502-521
179	481-500	502-521
180	481-500	502-521
181	481-500	502-521
182	481-500	502-521
183	481-500	502-520*
184	481-500	502-521
185	481-500	502-521
186	481-500	502-521
187	481-500	502-521
188	481-500	502-521
189	481-500	502-521
190	481-500	502-521
191	481-500	502-521
192	465-484	486-505
193	481-500	502-521
194	481-500	502-521
195	481-500	502-521
196	481-500	502-521
197	481-500	502-521
198	481-500	502-521
199	481-500	502-521
200	481-500	502-521
201	481-500	502-521
202	481-500	502-521

28/65

FIG. 6 (following)

203	479-498	500-519
204	481-500	502-521
205	481-500	502-521
206	481-500	502-521
207	481-500	502-521
208	481-500	502-521
209	481-500	502-521
210	481-500	502-521
211	481-500	502-521
212	481-500	502-521
213	481-500	502-521
214	481-500	502-521
215	481-500	502-521
216	479-498	500-519
217	481-500	502-521
218	482-500*	502-521
219	454-473	475-493*
220	481-500	502-521
221	480-499	501-520
222	481-500	502-521
223	481-500	502-521
224	481-500	502-521
225	481-500	502-521
226	481-500	502-521
227	483-500*	502-521
228	481-500	502-521
229	481-500	502-521
230	481-500	502-521
231	481-500	502-521
232	481-500	502-521
233	481-500	502-521
234	481-500	502-521
235	481-500	502-521
236	481-500	502-520*
237	482-500*	502-521
238	482-500*	502-521
239	481-500	502-521
240	481-500	502-520*
241	481-500	502-521
242	482-501	503-522
243	481-500	502-521
244	481-500	502-521
245	481-500	502-521
246	481-500	502-521
247	481-500	502-521
248	481-500	502-521
249	481-500	502-521
250	482-500*	502-521
251	464-483	485-503*
252	482-500*	502-521
253	481-500	502-521

29/65

FIG. 6 (following)

254	481-500	502-521
255	481-500	502-521
256	482-500*	502-521
257	482-500*	502-521
258	233-252	254-273
259	359-378	380-399
260	414-433	435-454
261	483-502	504-523
262	134-153	155-173*
263	186-204*	206-225
264	332-350*	352-371
265	482-500*	502-521
266	479-498	500-519
267	482-500*	502-521
268	481-500	502-521
269	482-500*	502-521
270	481-500	502-520*
271	481-500	502-520*
272	481-500	502-520*
273	481-500	502-521
274	481-500	502-520*
275	481-500	502-521
276	481-500	502-521
277	481-500	502-521
278	481-500	502-521
279	481-500	502-521
280	481-500	502-521
281	481-500	502-521
282	481-500	502-521
283	485-504	506-525
284	481-500	502-521
285	481-500	502-521
286	481-500	502-521
287	481-500	502-521
288	481-500	502-521
289	481-500	502-521
290	481-500	502-521
291	483-502	504-523
292	481-500	502-521
293	481-500	502-521
294	481-500	502-521
295	481-500	502-521
296	481-500	502-521
297	481-500	502-521
298	481-500	502-521
299	481-500	502-521
300	481-500	502-520*
301	481-500	502-521
302	481-500	502-521
303	481-500	502-521
304	482-500*	502-521



30/65

FIG. 6 (following)

305	481-500	502-521
306	481-500	502-521
307	481-500	502-521
308	481-500	502-521
309	481-500	502-520*
310	481-500	502-521
311	481-500	502-521
312	481-500	502-521
313	481-500	502-521
314	235-254	256-275
315	481-500	502-521
316	646-665	667-686
317	481-500	502-521
318	229-248	250-269
319	481-500	502-521
320	481-500	502-521
321	481-500	502-521
322	481-500	502-520*
323	481-500	502-520*
324	481-500	502-521
325	481-500	502-521
326	482-500*	502-521
327	481-500	502-521
328	481-500	502-521
329	481-500	502-520*
330	481-500	502-520*
331	481-500	502-521
332	481-500	502-521
333	481-500	502-521
334	481-500	502-521
335	481-500	502-521
336	481-500	502-520*
337	481-500	502-521
338	481-500	502-521
339	481-500	502-521
340	481-500	502-521
341	481-500	502-521
342	482-500*	502-520*
343	481-500	502-521
344	481-500	502-521
345	482-500*	502-520*
346	481-500	502-521
347	481-500	502-521
348	460-479	481-500
349	481-500	502-521
350	478-497	499-518
351	484-502*	504-523
352	484-502*	504-523
353	483-502	505-523*
354	484-502*	525-543*
355	483-502	504-523

31/65

FIG. 6 (following)

356	483-502	504-523
357	480-498*	500-519
358	480-498*	500-519
359	479-498	500-519
360	479-498	500-519
361	479-498	500-519
362	479-498	500-518*
363	479-498	500-519
364	479-498	500-519
365	479-498	500-519
366	412-431	433-452
367	368-387	389-408
368	483-502	504-523
369	484-502*	504-523
370	414-433	435-454
371	418-437	439-458
372	483-502	504-523
373	231-250	252-271
374	408-427	429-447*
375	481-500	502-521
376	278-297	299-318
377	367-385*	387-406
378	437-456	458-477
379	481-500	502-521
380	479-498	500-519
381	479-498	500-519
382	479-498	500-519
383	483-502	504-523
384	483-502	504-523
385	481-500	502-521
386	286-305	307-326
387	477-496	498-517
388	482-500*	502-521
389	481-500	502-521
390	481-500	502-521
391	481-500	502-521
392	224-243	245-263*
393	481-500	502-521
394	33-52	54-73
395	173-192	194-213
396	481-500	502-521
397	327-346	348-366*
398	483-502	504-523
399	221-239*	241-260
400	481-500	502-521
401	365-383*	385-404
402	483-502	504-523
403	483-502	504-523
404	483-502	504-523
405	483-502	504-523
406	484-502*	504-523

FIG. 6 (following)

407	479-498	500-518*
408	482-500*	502-521
409	482-500*	502-521
410	479-498	500-518*
411	481-500	502-520*
412	483-502	504-523
413	483-502	504-523
414	483-502	504-523
415	483-502	504-523
416	483-502	504-523
417	483-502	504-523
418	481-500	502-520*
419	481-500	502-521
420	479-498	500-518*
421	479-498	500-519
422	479-498	500-519
423	481-500	502-520*
424	481-500	502-520*
425	479-498	500-519
426	481-500	502-520*
427	483-502	504-522*
428	483-502	504-523
429	344-363	365-384
430	481-500	502-521
431	481-500	502-521
432	469-488	490-509
433	481-500	502-521
434	481-500	502-521
435	481-500	502-521
436	481-500	-
437	-	502-520*
438	481-500	-
439	481-500	-
440	481-500	-
441	481-500	-
442	481-500	-
443	481-500	-
444	482-500	-
445	481-500	-
446	481-500	-
447	481-500	-
448	481-500	-
449	481-500	-
450	481-500	-
451	481-500	-
452	481-500	-
453	481-500	-
454	481-500	-
455	481-500	-
456	481-500	-
457	481-500	-

33/65

FIG. 6 (following)

458	481-500	-
459	481-500	-
460	481-500	-
461	481-500	-
462	481-500	-
463	481-500	-
465	409-428	-
466	481-500	-
467	286-305	-
468	481-500	-
469	481-500	502-521
470	481-500	502-521
471	481-500	502-521
472	481-500	502-521
473	192-211	213-232
474	481-500	502-521
475	481-500	502-521
476	481-500	502-521
477	481-500	502-521
478	481-500	502-521
479	245-264	266-285
480	481-500	502-521
481	481-500	502-521
482	481-500	502-521
483	481-500	502-521
484	481-500	502-521

34/65

**Fig. 7**

SEQ ID NO.	POSITION RANGE OF AMPLIFICATION PRIMERS	COMPLEMENTARY POSITION RANGE OF AMPLIFICATION PRIMERS
1	362-380	792-812
2	310-327	751-771
3	304-321	745-765
4	82-99	540-557
5	308-325	830-847
6	304-321	803-823
7	131-150	561-580
8	287-304	805-825
9	284-303	716-734
10	394-413	826-844
11	270-289	704-721
12	444-462	874-893
13	73-91	577-596
14	139-158	634-652
15	372-392	808-826
16	429-449	865-883
17	233-252	693-712
18	122-141	582-601
19	298-317	772-792
20	296-315	770-790
21	200-217	679-696
22	442-459	921-938
23	228-245	760-777
24	378-396	911-928
25	203-221	736-753
26	37-55	570-587
27	222-241	655-675
28	436-455	880-900
29	476-493	945-962
30	266-283	735-752
31	278-295	613-632
32	418-436	823-842
33	216-235	646-665
34	91-109	510-528
35	137-153	586-604
36	137-153	586-604
37	206-225	727-746
38	400-419	856-876
40	146-165	588-607
41	62-81	504-523
42	210-230	591-610
43	307-326	797-817
44	277-296	767-787
45	1-20	402-421
46	3-21	404-422
47	1-18	355-372

35/65

FIG. 7 (following)

48	1-18	356-375
49	108-125	539-556
50	259-276	592-609
51	229-246	630-649
52	325-342	659-676
53	357-377	795-815
54	186-205	621-641
56	1-18	416-435
57	1-18	416-435
58	1-18	416-435
59	185-203	593-610
60	1-18	424-442
61	1-18	424-442
62	1-19	401-420
63	1-18	412-431
64	1-18	412-431
65	1-18	412-431
66	1-18	412-431
67	1-19	403-422
68	1-19	403-422
69	1-19	336-353
70	1-18	406-425
71	1-18	406-425
72	1-18	406-425
73	1-18	406-425
74	1-18	405-424
75	1-18	405-424
76	1-18	405-424
77	1-18	334-352
78	228-247	660-678
79	298-318	806-826
80	132-152	586-603
81	132-152	586-603
82	308-328	779-798
83	122-141	598-618
84	390-409	768-788
85	323-339	800-819
86	411-427	761-777
87	212-230	590-608
88	154-174	746-763
89	124-143	647-664
90	1-18	417-437
91	1-18	417-437
92	1-18	406-426
93	3-24	405-429
94	3-24	405-429
95	1-25	400-420
96	1-25	400-420
97	1-25	400-420
98	1-18	405-427
99	1-21	396-420

36/65

FIG. 7 (following)

100	1-21	396-420
101	1-18	423-443
102	1-18	423-443
103	1-18	429-446
104	1-18	429-446
105	1-18	429-446
106	1-25	399-420
107	1-23	398-418
108	1-23	398-418
109	1-18	427-445
110	1-18	427-445
111	1-18	416-436
112	1-18	416-436
113	1-25	396-420
114	1-25	396-420
115	1-25	396-420
116	1-18	416-434
117	1-21	396-420
118	1-19	405-429
119	1-19	405-429
120	1-21	422-439
121	1-21	422-439
122	1-21	422-439
123	1-21	422-439
124	1-21	422-439
125	1-21	413-432
126	1-18	404-423
127	1-21	411-435
128	1-21	411-435
129	1-18	428-449
130	1-18	395-419
131	3-23	398-420
132	1-21	418-442
133	1-21	418-442
134	1-21	418-442
135	1-21	418-442
136	1-24	404-426
137	1-24	404-426
138	1-24	404-426
139	1-24	404-426
140	1-20	396-420
141	459-476	859-878
142	126-143	526-545
143	451-468	853-872
144	388-407	758-775
145	226-245	705-725
146	220-238	636-655
147	478-496	820-837
148	95-112	504-521
149	297-317	742-759
150	416-435	868-886

37/65

FIG. 7 (following)

151	139-157	579-599
152	139-157	579-599
153	122-140	562-582
154	472-491	926-945
155	449-468	982-999
156	86-105	615-633
157	285-305	751-770
158	290-310	756-775
159	184-202	616-634
160	113-131	545-563
161	85-102	534-552
162	313-331	792-812
163	252-272	681-701
164	252-272	681-701
165	252-272	681-701
166	252-272	681-701
167	403-422	927-947
168	81-101	513-532
169	154-173	685-705
170	53-70	558-578
171	248-267	684-704
172	447-464	849-867
173	394-411	796-814
174	235-252	637-655
175	478-495	889-908
176	113-132	518-537
177	400-417	800-819
178	430-447	830-849
179	182-199	582-601
180	142-159	542-561
181	415-434	821-840
182	442-460	870-889
183	439-458	946-966
184	113-132	620-640
185	10-29	517-537
186	370-387	812-832
187	257-274	699-719
188	68-86	533-553
189	192-210	740-758
190	20-38	568-586
191	108-126	566-585
192	453-471	911-930
193	62-82	580-600
194	294-314	812-832
195	295-315	813-833
196	6-26	524-544
197	234-254	704-723
198	331-351	801-820
199	213-230	678-698
200	223-243	720-737
201	271-291	768-785



38/65

FIG. 7 (following)

202	468-488	965-982
203	311-331	802-820
204	86-106	528-546
205	189-209	621-641
206	266-286	777-795
207	90-109	514-534
208	127-144	571-591
209	336-355	784-802
210	230-249	678-696
211	78-97	526-544
212	283-303	711-731
213	168-188	636-655
214	80-100	535-552
215	90-110	545-562
216	446-463	993-1013
217	406-423	953-973
218	75-92	622-642
219	422-441	982-1001
220	410-429	970-990
221	341-360	901-921
222	324-343	884-904
223	473-491	907-925
224	460-479	889-909
225	376-395	805-825
226	107-127	539-559
227	125-145	558-575
228	191-208	596-613
229	382-400	805-824
230	326-345	728-747
231	148-167	550-569
232	240-257	658-675
233	276-293	692-711
234	136-155	581-598
235	424-444	855-875
236	348-368	784-803
237	105-125	541-560
238	437-456	839-859
239	436-455	838-858
240	384-402	832-849
241	75-94	544-563
242	76-95	545-564
243	280-299	749-768
244	346-365	815-834
245	364-383	833-852
246	363-381	878-893
247	171-189	686-707
248	159-177	674-695
249	17-35	532-553
250	369-386	777-794
251	237-253	567-586
252	221-238	624-641

39/65

FIG. 7 (following)

253	390-410	844-861
254	454-474	883-901
255	411-431	840-858
256	345-365	774-792
257	354-372	784-804
258	9-29	439-458
259	9-29	439-458
260	9-29	439-458
261	173-192	645-665
262	1-18	367-384
263	1-18	406-425
264	1-18	406-425
265	88-107	552-572
266	478-498	857-877
267	60-77	585-603
268	190-210	636-654
269	384-402	830-849
270	138-158	658-675
271	378-397	805-825
272	307-325	704-724
273	153-171	550-570
274	240-260	668-688
275	429-446	858-878
276	81-98	510-530
277	302-322	791-811
278	109-129	598-618
279	74-94	583-602
280	423-443	876-896
281	291-311	671-690
282	284-304	712-732
283	365-385	866-884
284	169-189	605-625
285	135-155	596-615
286	450-469	894-914
287	427-446	871-891
288	383-402	888-908
289	126-146	558-577
290	324-341	662-681
291	301-320	701-720
292	190-208	593-611
293	448-465	848-867
294	146-165	546-565
295	354-372	779-798
296	253-271	678-697
297	228-245	645-664
298	222-239	639-658
299	165-182	582-601
300	178-196	637-656
301	162-180	595-612
302	122-140	555-572
303	334-352	830-848

40/65

FIG. 7 (following)

304	309-327	805-823
305	203-221	699-717
306	444-462	940-958
307	267-286	703-722
308	342-361	778-797
309	276-295	706-725
310	326-344	779-797
311	301-319	733-753
312	244-262	676-696
313	104-124	594-612
314	99-117	557-577
315	68-86	526-546
316	250-270	800-818
317	250-270	800-818
318	12-32	442-461
319	186-204	623-641
320	87-105	524-542
321	112-130	549-567
322	122-139	602-620
323	368-386	868-888
324	173-191	673-693
325	117-135	617-637
326	181-200	640-658
327	414-433	873-891
328	187-205	637-657
329	181-199	631-651
330	50-68	533-552
331	148-167	651-669
332	12-31	515-533
333	271-291	655-674
334	144-164	528-547
335	375-395	765-783
336	403-423	793-811
337	368-387	800-820
338	328-345	827-845
339	306-323	805-823
340	232-249	731-749
341	50-68	553-572
342	386-405	867-887
343	23-43	522-540
344	99-116	628-647
345	412-429	844-862
346	67-87	513-533
347	230-247	691-710
348	446-463	907-926
349	345-363	729-749
350	478-497	909-927
351	479-498	880-899
352	420-439	788-807
353	420-439	788-807
354	304-322	714-732

41/65

FIG. 7 (following)

355	171-189	582-600
356	220-238	624-641
357	230-250	698-717
358	144-164	573-593
359	290-310	719-739
360	156-176	629-649
361	296-316	769-789
362	324-344	797-817
363	412-432	885-905
364	127-147	600-618
365	129-149	602-620
366	163-183	636-654
367	163-183	636-654
368	117-135	592-612
369	87-105	562-582
370	6-26	474-494
371	6-26	474-494
372	359-378	788-808
373	129-147	607-627
374	129-147	607-627
375	100-118	578-598
376	184-203	615-635
377	184-203	615-635
378	184-203	615-635
379	131-150	562-582
380	390-410	903-921
381	76-96	595-613
382	168-188	687-705
383	108-128	566-586
384	384-404	863-883
385	120-139	552-572
386	173-193	692-712
387	418-435	979-1000
388	353-372	809-829
389	348-367	804-824
390	251-270	707-727
391	52-71	508-528
392	46-65	533-553
393	20-39	507-527
394	18-38	505-525
395	18-38	505-525
396	18-38	505-525
397	272-292	704-724
398	209-229	641-661
399	219-237	653-673
400	437-457	908-928
401	279-298	773-793
402	252-271	746-766
403	238-257	660-679
404	172-189	578-597
405	135-152	545-564

42/65

FIG. 7 (following)

406	346-366	801-821
407	386-406	847-865
408	335-355	787-807
409	237-257	680-700
410	121-140	565-584
411	339-357	781-801
412	220-238	621-639
413	283-301	686-704
414	272-290	675-693
415	269-287	672-690
416	268-286	671-689
417	219-237	622-640
418	444-464	901-921
419	179-199	689-707
420	114-134	597-615
421	175-195	658-676
422	228-248	711-729
423	251-271	776-795
424	461-481	981-1001
425	343-363	781-799
426	424-442	952-971
427	309-326	777-797
428	455-473	907-927
429	40-59	551-569
430	114-134	562-582
431	115-135	563-583
432	127-147	575-595
433	179-199	627-647
434	321-341	769-789
435	326-346	774-794
436	367-385	797-817
437	184-203	616-633
438	86-106	546-563
439	260-279	755-773
440	255-272	773-790
441	303-322	736-756
442	215-233	617-635
443	479-499	913-932
444	204-222	634-654
445	104-121	506-525
446	370-387	769-793
447	475-495	870-892
448	347-367	764-788
449	111-132	513-537
450	96-113	601-621
451	149-167	577-596
452	312-329	713-731
453	128-145	530-548
454	444-461	855-874
455	76-95	583-603
456	333-353	774-793

FIG. 7 (following)

457	136-156	567-586
458	286-306	714-734
459	152-172	585-602
460	59-78	523-543
461	251-271	771-788
462	119-136	679-696
463	170-188	574-591
465	179-198	611-631
466	83-103	554-574
467	279-298	773-793
468	340-360	788-808
469	177-196	576-595
470	465-484	864-883
471	270-288	527-545
472	325-345	783-800
473	1-20	413-432
474	213-231	613-631
475	158-176	558-576
476	128-146	528-546
477	307-324	700-719
478	474-491	890-909
479	1-19	404-422
480	387-407	833-853
481	454-472	919-939
482	113-133	631-651
483	288-305	753-773
484	174-193	573-592

**Fig. 8**

SEQ ID NO.	POSITION RANGE OF PROBES
1	489-513
2	489-513
3	489-513
4	489-513
5	449-473
6	489-513
7	489-513
8	489-513
9	489-513
10	489-513
11	489-513
12	489-513
13	489-513
14	489-513
15	489-513
16	489-513
17	489-513
18	489-513
19	489-513
20	489-513
21	489-513
22	489-513
23	489-513
24	489-513
25	489-513
26	489-513
27	489-513
28	489-513
29	489-513
30	489-513
31	489-513
32	489-513
33	489-513
34	489-513
35	220-244
36	274-298
37	488-512
38	468-492
40	414-438
41	489-513
42	491-515
43	489-513
44	489-513
45	318-342
46	330-354
47	265-289
48	285-309

Fig. 8 (following)

49	489-513
50	489-513
51	489-513
52	489-513
53	489-513
54	489-513
56	166-190
57	206-230
58	322-346
59	489-513
60	66-90
61	92-116
62	40-64
63	27-51
64	119-143
65	154-178
66	234-258
67	149-173
68	150-174
69	157-181
70	140-164
71	231-255
72	265-289
73	283-307
74	31-55
75	105-129
76	308-332
77	109-133
78	489-513
79	438-462
80	234-258
81	299-323
82	487-511
83	489-513
84	489-513
85	491-515
86	489-513
87	491-515
88	489-513
89	489-513
90	204-228
91	385-409
92	220-244
93	39-63
94	114-138
95	190-214
96	263-287
97	334-358
98	159-183
99	176-200
100	211-235



46/65

**Fig. 8 (following)**

101	95-119
102	365-389
103	272-296
104	144-168
105	367-391
106	211-235
107	227-251
108	358-382
109	92-116
110	383-407
111	31-55
112	126-150
113	70-94
114	154-178
115	232-256
116	103-127
117	24-48
118	173-197
119	366-390
120	130-154
121	154-178
122	192-216
123	193-217
124	301-325
125	56-80
126	345-369
127	244-268
128	341-365
129	224-248
130	127-151
131	201-225
132	26-50
133	171-195
134	323-347
135	382-406
136	27-51
137	60-84
138	144-168
139	273-297
140	155-179
141	489-513
142	489-513
143	489-513
144	465-489
145	421-445
146	489-513
147	491-515
148	489-513
149	489-513
150	489-513
151	345-369

47/65

Fig. 8 (following)

152	461-485
153	489-513
154	489-513
155	467-491
156	488-512
157	489-513
158	489-513
159	491-515
160	491-515
161	489-513
162	489-513
163	369-393
164	449-473
165	461-485
166	479-503
167	489-513
168	489-513
169	489-513
170	489-513
171	489-513
172	489-513
173	489-513
174	489-513
175	485-509
176	489-513
177	489-513
178	489-513
179	489-513
180	489-513
181	489-513
182	489-513
183	489-513
184	489-513
185	489-513
186	489-513
187	489-513
188	489-513
189	489-513
190	489-513
191	489-513
192	473-497
193	489-513
194	489-513
195	489-513
196	489-513
197	489-513
198	489-513
199	489-513
200	489-513
201	489-513
202	489-513

48/65

**Fig. 8 (following)**

203	487-511
204	489-513
205	489-513
206	489-513
207	489-513
208	489-513
209	489-513
210	489-513
211	489-513
212	489-513
213	489-513
214	489-513
215	489-513
216	487-511
217	489-513
218	489-513
219	462-486
220	489-513
221	488-512
222	489-513
223	489-513
224	489-513
225	489-513
226	489-513
227	489-513
228	489-513
229	489-513
230	489-513
231	489-513
232	489-513
233	489-513
234	489-513
235	489-513
236	489-513
237	489-513
238	489-513
239	489-513
240	489-513
241	489-513
242	490-514
243	489-513
244	489-513
245	489-513
246	489-513
247	489-513
248	489-513
249	489-513
250	489-513
251	472-496
252	489-513
253	489-513

49/65

Fig. 8 (following)

254	489-513
255	489-513
256	489-513
257	489-513
258	241-265
259	367-391
260	422-446
261	491-515
262	142-166
263	193-217
264	339-363
265	489-513
266	487-511
267	489-513
268	489-513
269	489-513
270	489-513
271	489-513
272	489-513
273	489-513
274	489-513
275	489-513
276	489-513
277	489-513
278	489-513
279	489-513
280	489-513
281	489-513
282	489-513
283	493-517
284	489-513
285	489-513
286	489-513
287	489-513
288	489-513
289	489-513
290	489-513
291	491-515
292	489-513
293	489-513
294	489-513
295	489-513
296	489-513
297	489-513
298	489-513
299	489-513
300	489-513
301	489-513
302	489-513
303	489-513
304	489-513

Fig. 8 (following)

305	489-513
306	489-513
307	489-513
308	489-513
309	489-513
310	489-513
311	489-513
312	489-513
313	489-513
314	243-267
315	489-513
316	654-678
317	489-513
318	237-261
319	489-513
320	489-513
321	489-513
322	489-513
323	489-513
324	489-513
325	489-513
326	489-513
327	489-513
328	489-513
329	489-513
330	489-513
331	489-513
332	489-513
333	489-513
334	489-513
335	489-513
336	489-513
337	489-513
338	489-513
339	489-513
340	489-513
341	489-513
342	489-513
343	489-513
344	489-513
345	489-513
346	489-513
347	489-513
348	468-492
349	489-513
350	486-510
351	491-515
352	491-515
353	491-515
354	491-515
355	491-515

51/65

Fig. 8 (following)

356	491-515
357	487-511
358	487-511
359	487-511
360	487-511
361	487-511
362	487-511
363	487-511
364	487-511
365	487-511
366	420-444
367	376-400
368	491-515
369	491-515
370	422-446
371	426-450
372	491-515
373	239-263
374	416-440
375	489-513
376	286-310
377	374-398
378	445-469
379	489-513
380	487-511
381	487-511
382	487-511
383	491-515
384	491-515
385	489-513
386	294-318
387	485-509
388	489-513
389	489-513
390	489-513
391	489-513
392	232-256
393	489-513
394	41-65
395	181-205
396	489-513
397	335-359
398	491-515
399	228-252
400	489-513
401	372-396
402	491-515
403	491-515
404	491-515
405	491-515
406	491-515

52/65

Fig. 8 (following)

407	487-511
408	489-513
409	489-513
410	487-511
411	489-513
412	491-515
413	491-515
414	491-515
415	491-515
416	491-515
417	491-515
418	489-513
419	489-513
420	487-511
421	487-511
422	487-511
423	489-513
424	489-513
425	487-511
426	489-513
427	491-515
428	491-515
429	352-376
430	489-513
431	489-513
432	477-501
433	489-513
434	489-513
435	489-513
436	489-513
437	489-513
438	489-513
439	489-513
440	489-513
441	489-513
442	489-513
443	489-513
444	489-513
445	489-513
446	489-513
447	489-513
448	489-513
449	489-513
450	489-513
451	489-513
452	489-513
453	489-513
454	489-513
455	489-513
456	489-513
457	489-513

53/65

**Fig. 8 (following)**

458	489-513
459	489-513
460	489-513
461	489-513
462	489-513
463	489-513
464	489-513
465	417-441
466	489-513
467	294-318
468	489-513
469	489-513
470	489-513
471	489-513
472	489-513
473	200-224
474	489-513
475	489-513
476	489-513
477	489-513
478	489-513
479	253-277
480	489-513
481	489-513
482	489-513
483	489-513
484	489-513



54/65

## ALLELE FREQUENCY DATA (FRENCH and US)

Seq. ID No.	Protein	Biallelic Marker ID	caucasian FRENCH					caucasian US				
			size	A	C	G	T	size	A	C	G	T
21	MGST-II	12-458/198	no genotyped for this population					180	80,83			19,17
15	MGST-II	12-455/328						93	58,06		41,94	
5	MGST-II	12-426/154						181	58,01		41,99	
9	MGST-II	12-441/233						182		35,44		64,56
437	MGST-II	12-442/133						94		5,85	94,15	
7	MGST-II	12-430/80						93	5,38		94,62	
1	MGST-II	12-421/140						183	79,51		20,49	
436	MGST-II	12-421/139						185			84,59	15,41
13	MGST-II	12-453/429						183		56,28		43,72
12	MGST-II	12-447/58						93		55,38	44,62	
25	MGST-II	12-461/299						183		40,44		59,56
3	MGST-II	12-424/198						176			61,08	38,92
14	MGST-II	12-454/363						188	18,62		81,38	

Figure 9

55/65

## ALLELE FREQUENCY DATA (US)

Seq. ID No.	Protein	Blallelic Marker ID	caucasian US				
			size	A	C	G	T
37	ME1	12-716-295	189		66,14		33,86
31	ME1	10-428-219	189	66,67		33,33	
38	ME1	12-720-80	190		33,95	66,05	
33	ME1	10-420-284	184		1,36		98,64
41	ME1	12-721-440	94	1,6		98,4	
44	ME1	12-724-225	190		73,16		26,84
42	ME1	12-723-293	92		99,46		0,54
35	ME1	12-713-95	190		32,63		67,37

Figure 10

56/65

## ALLELE FREQUENCY DATA (FRENCH and US)

Seq. ID No.	Protein	Biallelic Marker ID	Caucasian FRENCH					Caucasian US				
			size	A	C	G	T	size	A	C	G	T
336	UGT1A7	12-148-311	88	47,73		52,27		190	44,47		55,53	
345	UGT1A7	12-156-91	90	50,56		49,44		93	50,54		49,46	
300	UGT1A7	12-121-326	93	30,11		69,89		188	28,99		71,01	
309	UGT1A7	12-128-225	93	50,54	49,46			187	55,35	44,65		
304	UGT1A7	12-124-194	85		19,41		80,59					
326	UGT1A7	12-141-159	92	37,5		62,5						
330	UGT1A7	12-143-453	82	50,61		49,39						
322	UGT1A7	12-139-380	85	17,65		82,35		190	20,79		79,21	
342	UGT1A7	12-153-116	90		62,22		37,78					
323	UGT1A7	12-140-134	89			58,99	41,01	183			56,01	43,99
329	UGT1A7	12-142-321	90	54,44		45,56						

Figure 11

57/65

# ALLELE FREQUENCY DATA (FRENCH and US)

Seq. ID No.	Protein	Biallelic Marker ID	Caucasian FRENCH					Caucasian US				
			size	A	C	G	T	size	A	C	G	T
380	UGT2B4	12-653-423	6	25			75	177	73,45			26,55
351	UGT2B4	10-470-25	179	56,15			43,85	90	55,56			44,44
356	UGT2B4	10-494-284	6		16,67		83,33	185		22,97		77,03
352	UGT2B4	10-471-84	178	24,44			75,56	92	27,17			72,83
353	UGT2B4	10-471-85	0					188	26,6	73,4		
354	UGT2B4	10-472-202	182		14,29		85,71	93		9,68		90,32
357	UGT2B4	12-637-219	0					187		33,42		66,58
358	UGT2B4	12-639-95	0					187		35,56		64,44
377	UGT2B4	12-652-203	4	50	50			189	57,67	42,33		
369	UGT2B4	12-642-417	0					186	73,66		26,34	
374	UGT2B4	12-648-300	0					91		62,09		37,91

Figure 12

### **297 ALT vs 286 Caucasian US**

58/65

### Figure 13

## HAPLOTYPE ANALYSIS : PERMUTATION TEST RESULTS

Asthma

HAPLOTYPE GCT	sample sizes cases vs controls	markers in bac										MARKERS	
		G					T					HAPLOTYPES (ALT vs US)	
		C					T					HAPLOTYPES (ALT vs US)	
		G					T					HAPLOTYPES (ALT vs US)	
ALT+ vs ALT-	60 vs 81	0.077	0.141	0.077	0.141	0.077	0.141	0.077	0.141	0.077	0.141	pvalue	ALT+ vs ALT-
ALT vs caucasian US	151 vs 86	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	diff all. Freq%	ALT vs caucasian US
ALT vs caucasian US (2)	151 vs 88	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	diff all. Freq%	ALT vs caucasian US
HAPLOTYPE GCTG	sample sizes cases vs controls	G					T					MARKERS	
		C					T					HAPLOTYPES (ALT vs US)	
		G					T					HAPLOTYPES (ALT vs US)	
		G					T					HAPLOTYPES (ALT vs US)	
ALT+ vs ALT-	60 vs 81	0.077	0.141	0.077	0.141	0.077	0.141	0.077	0.141	0.077	0.141	pvalue	ALT+ vs ALT-
ALT vs caucasian US	151 vs 86	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	diff all. Freq%	ALT vs caucasian US
ALT vs caucasian US (2)	151 vs 88	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	0.118	0.011	diff all. Freq%	ALT vs caucasian US
HAPLOTYPE CATT	sample sizes cases vs controls	G					T					MARKERS	
		C					T					HAPLOTYPES (ALT vs US)	
		G					T					HAPLOTYPES (ALT vs US)	
		G					T					HAPLOTYPES (ALT vs US)	
ALT+ vs ALT-	64 vs 102	0.139	0.075	0.139	0.075	0.139	0.075	0.139	0.075	0.139	0.075	pvalue	ALT+ vs ALT-
ALT vs caucasian US	166 vs 92	0.101	0	0.101	0	0.101	0	0.101	0	0.101	0	diff all. Freq%	ALT vs caucasian US
ALT vs caucasian US (2)	166 vs 92	0.101	0	0.101	0	0.101	0	0.101	0	0.101	0	diff all. Freq%	ALT vs caucasian US

Figure 14

89 ALT+ vs 208 ALT-

MARKERS		ESTIMATED FREQUENCIES													
MGST2		markers in bac													
		12-471/136	12-471/140	12-430/80	12-441/233	12-442/133	12-447/58	12-485/226	12-461/299	12-453/429	12-424/198	12-454/303	12-458/196	12-426/154	
		inB													
		88 vs 194	88 vs 195	68 vs 97	87 vs 198	68 vs 104	89 vs 198	65 vs 102	88 vs 199	88 vs 199	80 vs 180	83 vs 194	83 vs 194	85 vs 188	
		18/12 (T)	18/18 (G)	91/81 (G)	39/34 (C)	92/81 (G)	47/44 (G)	56/55 (A)	64/62 (T)	40/38 (T)	41/40 (T)	22/20 (A)	22/16 (T)	60/59 (A)	
		5.6	0.4	0.4	5.6	1.3	3.1	0.8	1.8	2.7	1.0	2.5	5.5	1.8	
		7.8e-02	7.5e-01	7.5e-01	1.9e-01	6.6e-01	4.8e-01	7.5e-01	6.6e-01	5.3e-01	7.5e-01	4.8e-01	1.2e-01	6.6e-01	
		1.5	1.0	1.1	1.3	1.2	1.1	1.0	1.1	1.1	1.0	1.2	1.4	1.1	
		-0.01 HW	0.00 HW	0.83	0.02 HW	0.85	-0.04 HW	-0.02 HW	0.01 HW	-0.01 HW	0.04 HW	-0.00 HW	0.03 HW	-0.00 HW	
		0.00 HW	-0.00 HW	0.00 HW	0.01 HW	0.00 HW	0.01 HW	0.03 HW	0.01 HW	-0.03 HW	0.01 HW	0.00 HW	-0.00 HW	0.03 HW	
</															

**Fig. 15**

PROTEIN MGST2	12-441/233	12-461/299	12-453/429	12-426/154	MARKERS	
	markers in bac					
	C		T		HAPLOTYPES (ALT+ vs ALT-)	
					pvalue	
	1.92E-01	6.55E-01	5.27E-01			
	-5.6 (40 vs 34)	1.6 (64 vs 63)	-2.7 (41 vs 38)			ALT+ vs ALT-
	7.52E-01	2.73E-01	1.47E-01			
	-0.3 (36 vs 35)	3.5 (63 vs 60)	4.7 (39 vs 44)			ALT vs caucasian US
	C		T		HAPLOTYPES (ALT+ vs ALT-)	
	1.92E-01	6.55E-01	5.27E-01	6.55E-01		
	-5.6 (40 vs 34)	1.6 (64 vs 63)	-2.7 (41 vs 38)	1.8 (61 vs 59)		ALT+ vs ALT-
	7.52E-01	2.73E-01	1.47E-01	6.55E-01		
	-0.3 (36 vs 35)	3.5 (63 vs 60)	4.7 (39 vs 44)	1.3 (59 vs 56)		ALT vs caucasian US

HAPLOTYPE CIT	sample sizes cases vs controls	haplotype frequencies		p-excess	odds- ratio	chi-S	P value	PERMUTATIONS		
		cases	controls					Av. Chi-S	Max Chi-S	> 1ter / nb of 1ter.
ALT+ vs ALT-	86 vs 198	0.157	0.049	11.34	3.63	18.68	1.5e-05	1.9	18.9	1/1000
ALT+ vs ALT- (1)	86 vs 104	0.157	0.088	7.54	1.93	4.26	3.8e-02	1.1	25.4	12/10000
ALT+ vs ALT- (2)	86 vs 94	0.157	0.042	11.91	4.19	13.36	2.5e-04	1.5	16.5	52/1000
ALT vs caucasian US	284 vs 176	0.092	0.071	2.2	1.32	1.18	2.7e-01	2.1	20.9	3/1000
ALT vs caucasian US (2)	284 vs 94	0.092	0.047	4.67	2.04	3.77	5.1e-02	1.9	17.3	158/1000
ALT vs caucasian US (3)	284 vs 82	0.092	0.116	-2.79	0.77	0.88	3.4e-01	1.8	17.7	498/1000

HAPLOTYPE CTTA	sample sizes cases vs controls	haplotype frequencies		p-excess	odds- ratio	chi-S	P value	PERMUTATIONS			
		cases	controls					Av. Chi-S	Max Chi-S	> iter /	
ALT+ vs ALT- ALT+ vs ALT- (1) ALT+ vs ALT- (2)	83 vs 187	0.141	0.028	11.64	5.75	25.13	5.2e-07	*****	2.6	25.2	1/1000
	83 vs 98	0.141	0.066	8.03	2.33	5.61	1.7e-02	**	2.5	35	7/10000
	83 vs 89	0.141	0.034	11.03	4.6	12.42	4.1e-04	***	1.7	17.4	59/1000
ALT vs caucasian US ALT vs caucasian US (2) ALT vs caucasian US (3)	270 vs 167	0.082	0.058	2.61	1.46	1.85	1.7e-01	*	1.7	18.3	3/1000
	270 vs 89	0.082	0.023	6.12	3.89	7.61	5.5e-03	**	2.7	25.9	438/1000
	270 vs 78	0.082	0.110	-3.06	0.73	1.11	2.7e-01	*	2.3	17.8	499/1000

### Figure 16



62/65

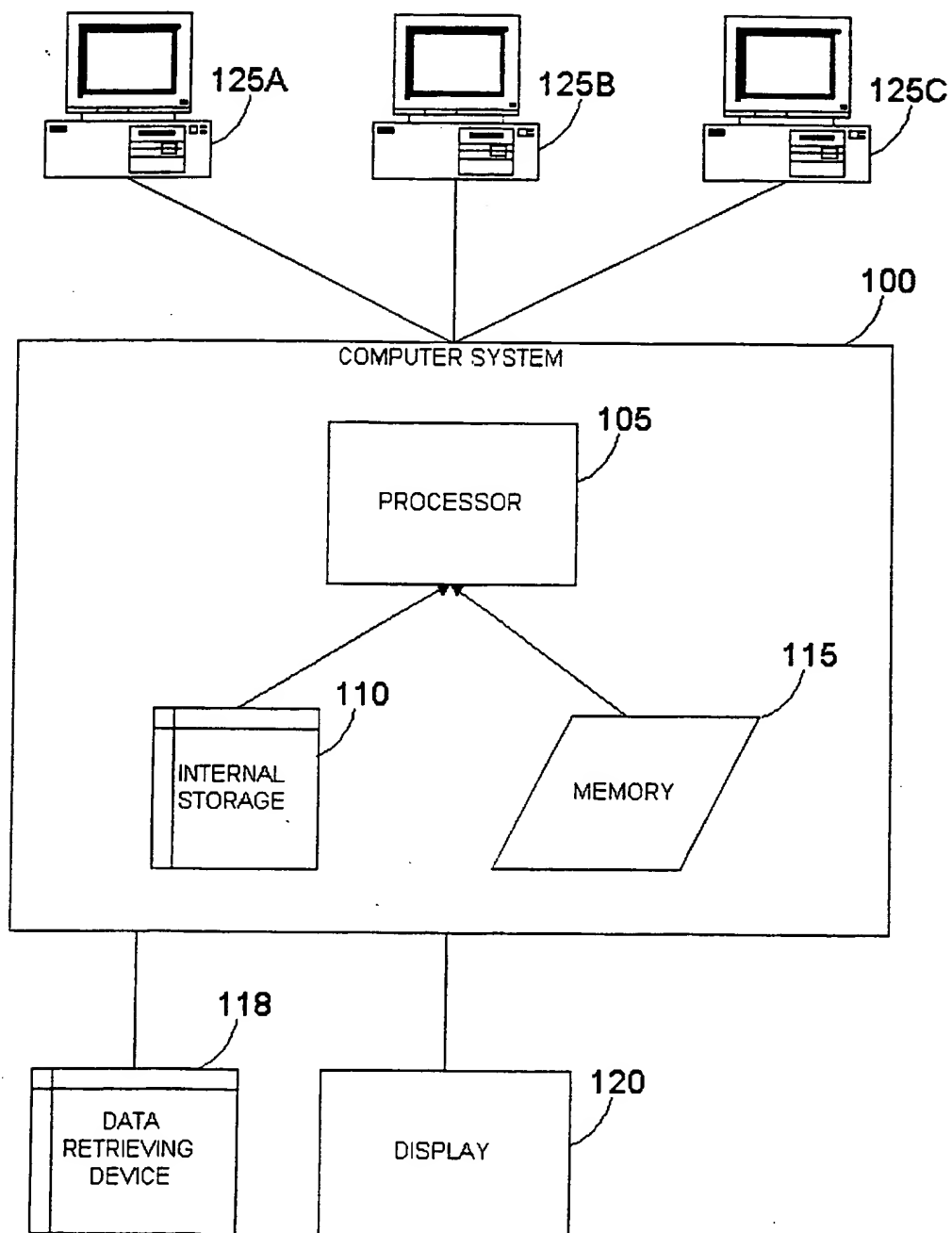


FIG.17

63/65

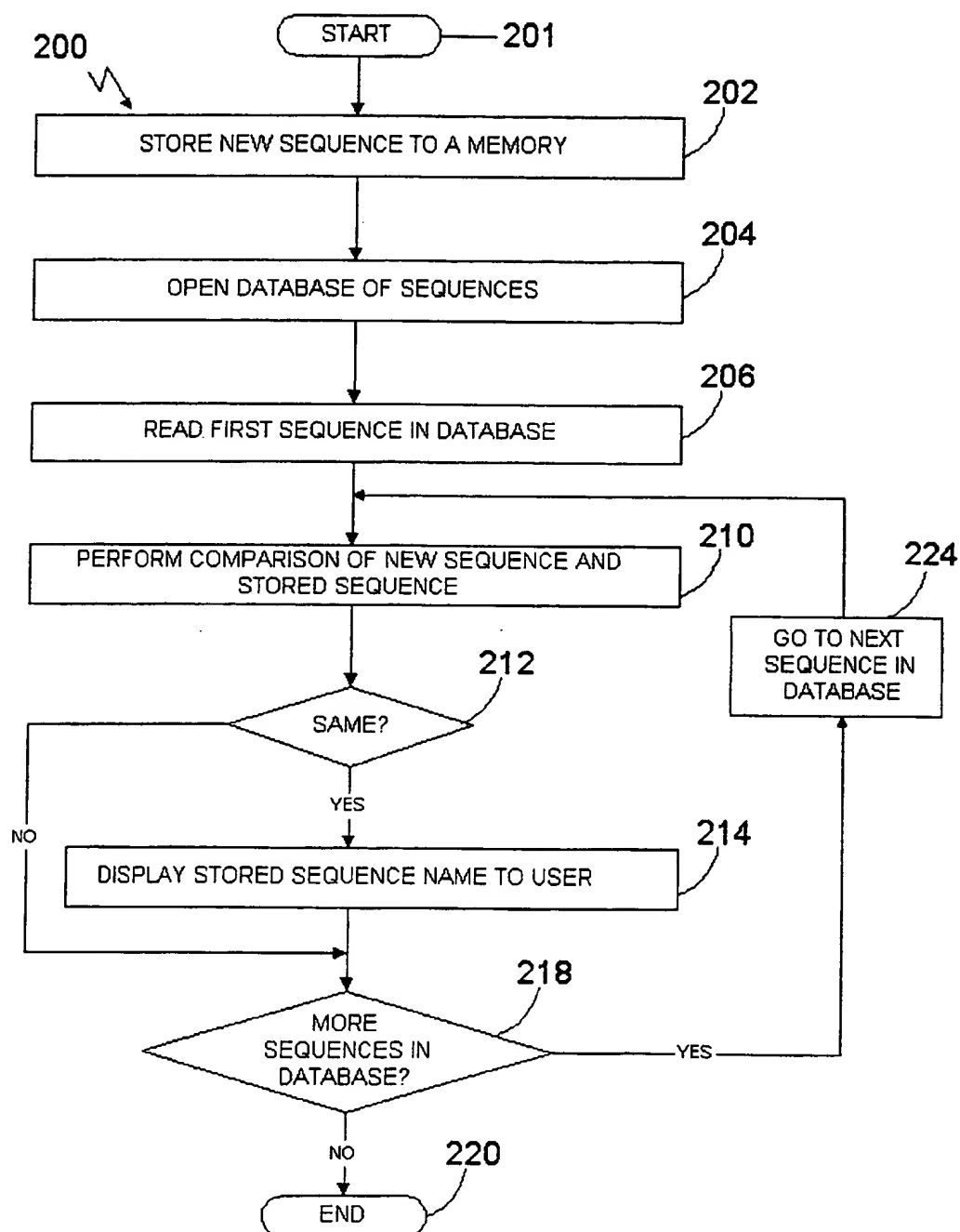


FIG.18

64/65

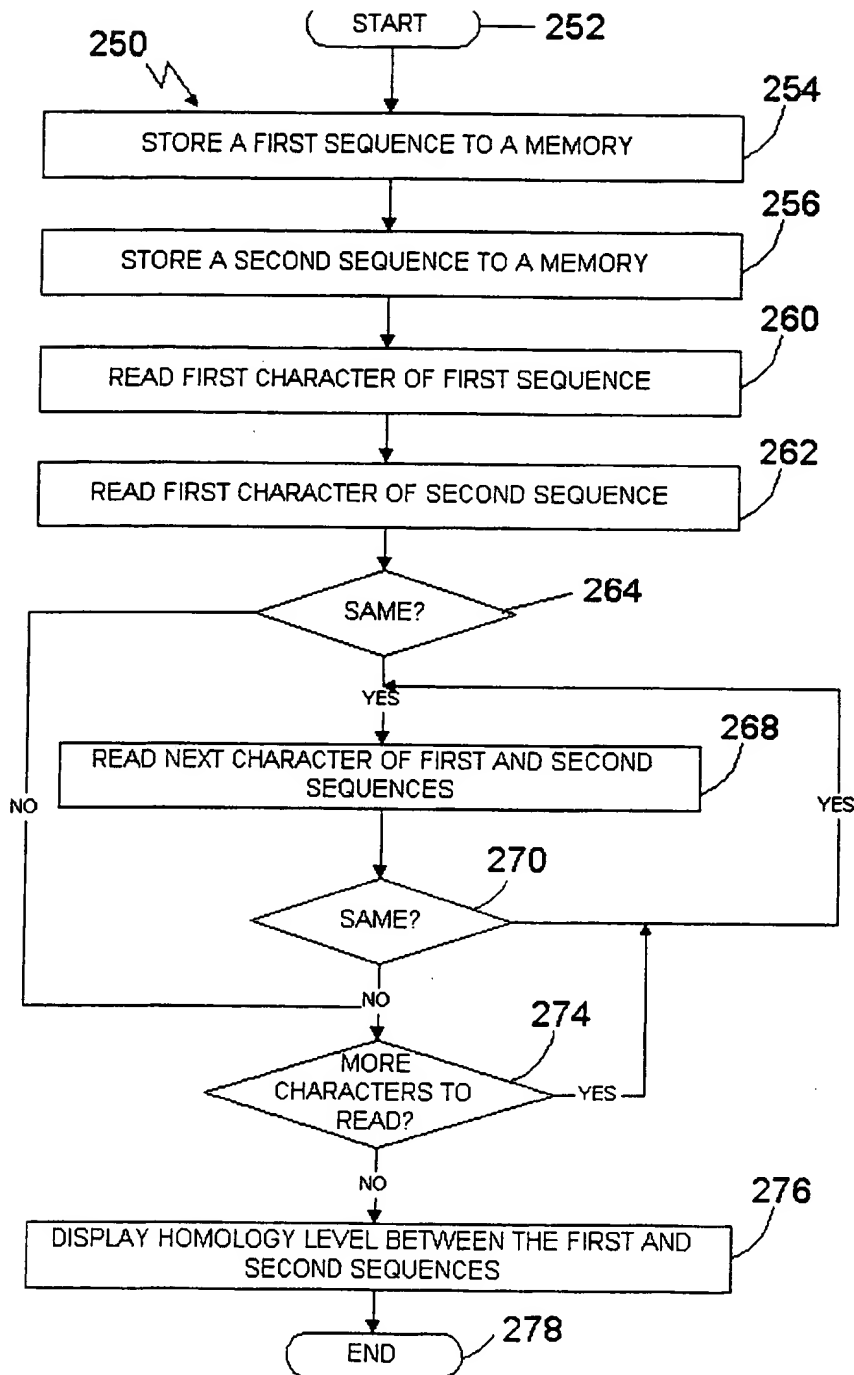


FIG.19

65/65

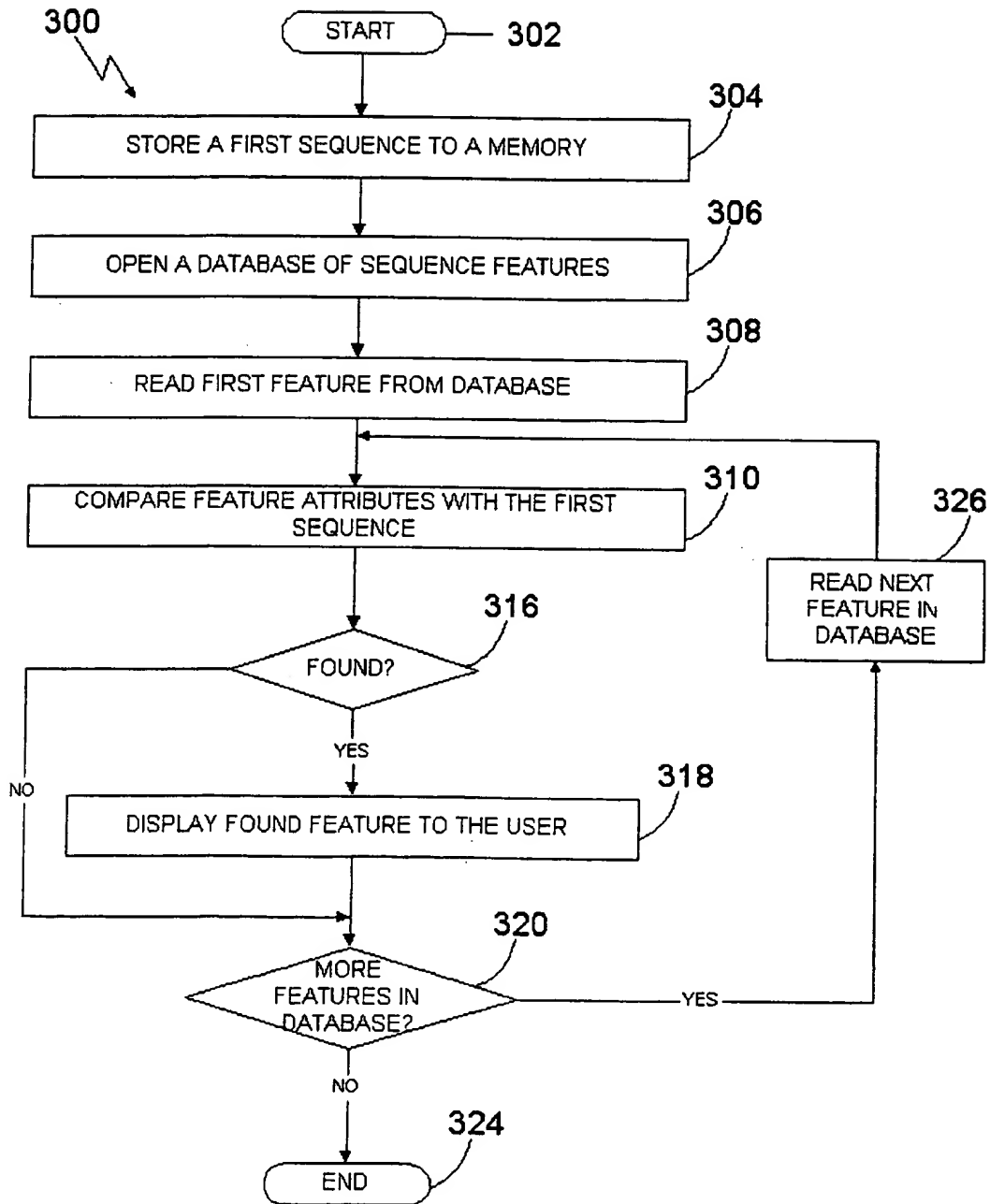


FIG.20

## SEQUENCE LISTING

&lt;110&gt; GENSET

&lt;120&gt; BIALLELIC MARKERS RELATED TO GENES INVOLVED IN DRUG METABOLISM

&lt;130&gt; 62.W01

&lt;150&gt; US 60/126,269

&lt;151&gt; 1999-03-25

&lt;150&gt; US 60/131,961

&lt;151&gt; 1999-04-30

&lt;160&gt; 493

&lt;170&gt; Patent.pm

&lt;210&gt; 1

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-421-140 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-421-140.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-421-140.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 362..380

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 792..812

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-421-140 potential probe

&lt;400&gt; 1

tgtctcattt	tgatacagat	gaacctagtg	tcaaatgagt	tggaggaagc	atcctgttct	60
actttctcag	ataattttaa	taaggttgga	agagtgtgtt	ccttcaagg	tgtgtagaac	120
ttgccttaaa	ataatctggg	cctaattgtt	ttctgggtgg	aaatttttaa	attatgggat	180
ttgcattctt	taatgattat	tggactaggc	aggctttttg	ttacatcttg	aatcaatttt	240
ggtaggcctt	ggctccttgag	atgataacat	tggtctcttg	attatccact	tgctccaagg	300
aggggttttac	agactactcc	ctaggcttga	aagcgtgggg	tgaataccac	cgggagtaca	360
cgagataatt	gtaagaggtg	tgtagacttg	gttttaaata	acattgaact	acatgatgag	420
aagtcattct	catttcaatt	atctttgaat	ccttatgggt	aagtcaagtg	gaaagtattg	480

2

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tgccccctacc tccccctctcc ctctatccct agagataacc ctttttgatt actactttgg 660
gtattttactt ctgtattttct aaacattata ctttctactgc tattttattga ttcacaaatt 720
ttagacattg tcagttggct tctttttatg gaagaaataa tttcatgctc ctttacttct 780
tcttttcctt acctcatcct tccagtaaca ttatatcact ctggttaggtt aaatcaatag 840
ccaatgttta taattattgt gattaagaaa ataattgtga aagcagggcc ttatagtata 900
ctgtgggttac atttccttct tgtgcaattt tttttcccta gggtaataa ttgctcttta 960
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&lt;211&gt; 999

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-424-192 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-424-192.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-424-192.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 310..327

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 751..771

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-424-192 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 521,541,635

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 2

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gtgggggggga tggggaattc ctctcttacc aaccctgcct catttcagtt tttgcttcag 180
aagcttgggc ttctcaccac ttctccatc tggctagctg gtctactcta agccctactg 240
tggggccact catccagggc tcacagaggc ccccatggta aggccaccag ctcttgattc 300
agcttccgca caatgtggga acttaggaca gctcaggaaa cctaactcag ggccttcctt 360
ctccttacca tactacttta ctctgctatg gctgtgtaaa agactgaatg ggacagagat 420
cttgtaggag gagagaaacg cccattagta atttccaagc atgcacacat gttatagggt 480
ctgaggggctc aggaaacagtt rcatgaggtc cagagttcga nttatgaagc tgaggaattt 540
nactaatgta ttcatctgtt gaatatttca gtgtattttg caagttggca ctgtgtgagt 600
gtcacacttg gctctacagt ggtagcaaa tctgnacaca gtcctcatgg atcttacata 660
agaggaaaaa actgacaacc aaatgatcac acaaatgagt aactacacag tatgcaatgc 720

```

3

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tatttggtga	atgaatgaat	aaataaatgc	atgaaagcta	tgagattact	tctgagaggt	900
cagggatgaa	ttcctgaggc	agtgttattt	tggctgatgt	taaagagaaa	aaaaaattct	960
tagttcgaat	tctgctagat	aaatttatat	taaccaaac			999

&lt;210&gt; 3

&lt;211&gt; 993

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-424-198 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-424-198.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-424-198.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 304..321

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 745..765

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-424-198 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 515,535,629

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 3

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aaactggcaa	ctgctagttc	agcaactcca	aggcaggcaa	tgcagaaaca	tggcgtgggg	120
gggatgggga	attcctctct	taccaaccct	gcctcatttc	agtttttgct	tcagaagctt	180
gggctttctca	ccactttctc	catctggeta	gctgggtctac	tctaagccct	actgtggggc	240
cactcatcca	gggctcacag	agggccccc	ggtaaggcca	ccagctcttg	attcagcttc	300
cgcacaatgt	gggaacttag	gacagctcag	gaaacctaac	tcagggcctt	ccttctccct	360
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ccaggagaga	aacgcccatt	agtaatttcc	aagcatgcac	acatgttata	ggttctgagg	480
gctcaggaac	agttgcatga	kgtccagagt	tcganttatg	aagctgagga	atttnactaa	540
tgtattcatt	cgttgaatat	ttcagtgtat	tttgcaagtt	ggcactgtgt	gagtgtctaca	600
cttggtctta	cagtggtttag	caaatctgna	cacagtcctc	atggatctta	cataagagga	660
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aaaggagggg	ctggttgcta	tcttcacgag	aataataagca	ccacagtggg	tttctttatt	780
ttgttcattg	atgtacctag	aatggggcct	cacacataatt	aagtgtctaaa	taagtatttg	840
ttgaatgaat	gaataaataa	atgcatgaaa	gctatgagat	tacttctgag	aggtcagggg	900
tgaattcctg	aggcagtgtt	attttggtg	atgttaaaga	gaaaaaaaaa	ttcttagttc	960

4

gaattctgct agataaattt atattaacca aac

993

&lt;210&gt; 4

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-425-57 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-425-57.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-425-57.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 540..557

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 82..99

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-425-57 potential probe

&lt;400&gt; 4

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aaactattat tatcttctcg cttagcggga ctgaaggata gcaagggtgct tggaaatttcc	180
cttgaaggaa ctcaagagtt tttatttccg tgcttgaggt ggctagcagg tccctaagag	240
tgttccttgc tctgtctcca tgccagaagt ttattatggg gtgaaaagag gaagttaagg	300
acaggaggca agacctactg aaggctcaaa aactgacaga ggaggaagta tggatgtgga	360
agtgaagaa gtccacagag ctcatcacia gatgaagggt gtcccaggca tcagagttga	420
ttatctaatag aaataaagga gaagaaagtc atactgtttt ataacaggga aacgtcggca	480
ggtaggagcc atcatcccca ragttttctt gtacctgctg ggcaagggtg cagagccctg	540
gcaatggaaa gtgattggcc cacaggaggt aagaatttac cgacaacaat ataggtttga	600
aaaggaaaagt tttattagat tgagagaaag ctgcagaaga gtgcagcggg ctgcctcagc	660
gagaggactg agcgcgccgg ggaggatttt ccttaggggt atttatggac cttaaagcaa	720
gagcttaggg gtaatttggt ccatattagc cacatatgtc atgataaatg attacatttg	780
tagacatttt ggtgtcgtaa tgtcagcaag gggtgcacaa tgagttttgt catgcatgca	840
ttccggagat gtatagaggt agagaaagtc tagttacata taaatttttg ggaaagaagc	900
ctggaaccgg ccttagatat agggaagatc aattatttct aaactcctca gataaggagt	960
tctgcctctg gatggtcggc ttcattggcca ccagggtgtc t	1001

&lt;210&gt; 5

&lt;211&gt; 961

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele



5

<222> 461  
 <223> 12-426-154 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 441..460  
 <223> 12-426-154.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 462..481  
 <223> 12-426-154.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 308..325  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 830..847  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 449..473  
 <223> 12-426-154 potential probe

<220>  
 <221> misc\_feature  
 <222> 705,761..762  
 <223> n=a, g, c or t

<400> 5  
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 ttcagtgcag gctaaagtgt gagaaactaca gatacaaaagg agggctctgt atctgcaaca 180  
 acaactatgt aatgatgact agtccccact cttctgaagc taccatggc aaaattagaa 240  
 acataattct tcaaaaacca ctggcactga tttagtgcac agaacacctg tggataactc 300  
 aattcccttt gtccctccct gtgtcaaggc caaaggatct aatgtgatat aagcattcca 360  
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 t 961

<210> 6  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-429-198 : polymorphic base C or T  
 <220>

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<221> misc_binding
<222> 481..500
<223> 12-429-198.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-429-198.mis2, potential complement

<220>
<221> primer_bind
<222> 304..321
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 803..823
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-429-198 potential probe

<220>
<221> misc_feature
<222> 44,561..562,954
<223> n=a, g, c or t

<400> 6
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caggagaatc gcttgaaccc aggaggcgga ggttgcagtg agttgacagt gtaccactgc      180
actccagcat gggcgacaga ccaagattct gtcttaaaaa acaaaacaaa acaaaacaaa      240
aaactgggag gggaaaatat ttttttattt tctaatatgt tagaaccctt tctcaatgtg      300
taactctcct cattcttctc cctctcatct ttaacaaaaa ttgtaaatgt tctacatgtg      360
catcatgtta gtctatcagg tgttatttag ataattggtg cctttgactt acattcttgt      420
aaacgttgct ttagttcata aactcataat attcccagac ctgaagcctg tcttaaaatt      480
gataatgtaa atcatctagg yctgaatgcc taaagtgatg gggagctcac cacctcttgt      540
ttctggtcat aggtagggtga nngatccctt ctgcagcaag aagtcagggg caggcctgtg      600
acactagctc tgattcctgc aaatgaccaa atgaggggca gggcctggtg gtctcccagt      660
ccagctgtag gctcccatga cctgcacctc cctgccctcc caactctcag gttctcccaa      720
cagccctccc ccagctaatt tcataaaagg cactaaaaac ttttgatcag tcacaaaatc      780
atgggaagag aaagaaatac aagaagagag aaagaaggga aagtggcgtg aatcattaac      840
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<210> 7
<211> 996
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-430-80 : polymorphic base T or C

<220>
<221> misc_binding
<222> 502..521
<223> 12-430-80.mis1, potential complement

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<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-430-80.mis2

<220>  
 <221> primer\_bind  
 <222> 561..580  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 131..150  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-430-80 potential probe

<220>  
 <221> misc\_feature  
 <222> 918,971,978  
 <223> n=a, g, c or t

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 acatctaatt ggtaagtgt taggtattcg ttgtattatt atttctactt tcctgggtgt 180  
 ttaaaaatgt ttataataaa gagttaaagg atctgaagag aagttaaaac ctaaaatcag 240  
 attcttggtg gccctgaaga ttttctgtgt tgagcattgt taagagatcc agtgggtattg 300  
 atggatccac agtcatccat gctggcctgg cagctgtcct ggggtgtggca cagatgcctg 360  
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 aatacgtctt tctcactggg ggcagatgtc tagtgacttt ctagaaaaat gtttattctc 840  
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<210> 8  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-433-215 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-433-215.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-433-215.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 287..304  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 805..825  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-433-215 potential probe

<220>  
 <221> misc\_feature  
 <222> 92,202  
 <223> n=a, g, c or t

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 caacctcatt gcaaggtaac aactcagctt cccagggttc atcatttggc tccagtata 360  
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 ctggtggccc ttaggtggca ggattgcaga ggccccaccg gaagtcagct ggccacagca 480  
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<210> 9  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-441-233 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-441-233.mis1, potential complement

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-441-233.mis2

<220>  
 <221> primer\_bind  
 <222> 716..734

<223> upstream amplification primer, complement

<220>

<221> primer\_bind

<222> 284..303

<223> downstream amplification primer

<220>

<221> misc\_binding

<222> 489..513

<223> 12-441-233 potential probe

<220>

<221> misc\_feature

<222> 661,929,990

<223> n=a, g, c or t

<400> 9

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aagcaattgc	tccaccttac	aaatttggat	agaaaggaga	tgtagtttat	ttcatatggg	360
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agtggatcaa	agagatgaaa	ccaaaaaana	agcaaacaaa	caaatgagaa	gacacaaaac	960
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<210> 10

<211> 1001

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 501

<223> 12-441-343 : polymorphic base G or A

<220>

<221> misc\_binding

<222> 481..500

<223> 12-441-343.mis1, potential

<220>

<221> misc\_binding

<222> 502..521

<223> 12-441-343.mis2, potential complement

<220>

<221> primer\_bind

<222> 826..844

<223> upstream amplification primer, complement

<220>

<221> primer\_bind

<222> 394..413  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-441-343 potential probe

<220>  
 <221> misc\_feature  
 <222> 3,771  
 <223> n=a, g, c or t

<400> 10  
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 ctcccttttg gcgaaatcct gggagtgatg ctctaaaaat ccacctttcc catcatccct 360  
 actcatcaga aagacaaata taaaatccca gagagggtga ggagctaaaa aagcaattgc 420  
 tocaccttac aaatttggat agaaaggaga tgtagtttat ttcatatggg caaagtagtc 480  
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<210> 11  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-442-221 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-442-221.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-442-221.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 704..721  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 270..289  
 <223> downstream amplification primer

<220>

<221> misc\_binding  
 <222> 489..513  
 <223> 12-442-221 potential probe

<220>  
 <221> misc\_feature  
 <222> 180,531..532,822  
 <223> n=a, g, c or t

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 agaggacata aacactgatt ttttcccccc attcatttaa ctatttgcac acagagacan 180  
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 caagccgcat cataaaggaa aagtatttct ttttgttctg gccaaagcaa aatacgcgta 360  
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 tttgccctca ggtgggccct gtcattctta atctaacctc cgactaggag tttcaacatg 480  
 tgggtctctgg gcaagatggt ygccctgagt aatagaaaag aaagagaaaag nngagagaga 540  
 gaaaaacatt gcctgtggca gggcggggaa ggtgaaatga tcaggaggagc agagaaagaa 600  
 ccaccattg cagcgacact aaaaagtcca ggtggctgct gtcggtggag caaggatctt 660  
 ttccagtaat ctaccagct ctcaaatttc cttgttagg gaggaaaaag ctcccatgt 720  
 cccaggatcc tgtacattcc taattctgtc acccatagcc atcagcaaag tacaagggag 780  
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 agggacttta ccgaagaggg gcctctaacc cgctaaatct tagaagggac tctaactctc 900  
 ctaagtcggg cctctaacca gaggtcagtc aagcatcctt gccttttatt aagggggagc 960  
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<210> 12  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-447-58 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-447-58.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-447-58.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 444..462  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 874..893  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-447-58 potential probe

12

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 44,894

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 12

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gaagtttggc	tgtttaaagg	gctattttta	atataggcaa	cagaaaacaa	aatacaaagt	840
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tatccccacc	tgtcataaaa	tggcaaatag	gctgggtgca	gtggctcatg	cctgtaatcc	960
cagcactttg	ggaggtcaag	ttgggaggat	catttgagtc	c		1001

&lt;210&gt; 13

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-453-429 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-453-429.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-453-429.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 73..91

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 577..596

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-453-429 potential probe

&lt;400&gt; 13

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aaggagcata	aacacaatca	caaaattggg	gctccaccca	tcggttgca	aggcagtcgg	120
cagtgtctgc	tacaagtgc	atcaagtagc	atctatcaag	tggcattact	ttacaaaata	180



13

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cagtttttat	tttttagtct	tggttggggt	cactgaaact	tccaggttgt	cctaacctgt	360
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acaaaaaaca	gggtagaagg	taataacaca	tgcaaaggga	tgtgtagcaa	tcaaagggaa	480
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&lt;210&gt; 14

&lt;211&gt; 929

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-454-363 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-454-363.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-454-363.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 139..158

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 634..652

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-454-363 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 674..679,881..882,892..893

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 14

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14

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&lt;210&gt; 15

&lt;211&gt; 827

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-455-326 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-455-326.mis1, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-455-326.mis2, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 808..826

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 372..392

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-455-326 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 798..799

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 15

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15

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<210> 16  
<211> 884  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-455-383 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-455-383.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-455-383.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 865..883  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 429..449  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-455-383 potential probe

<220>  
<221> misc\_feature  
<222> 855..856  
<223> n=a, g, c or t

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agaggaaatt gacaggaaga ggaattgaga agaattctat agaattgggt atgaacaatg 180  
gatcaggaac aatagcagaa taataaggcc tgcagaacta cctggaaacc tgaactctgg 240  
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aattttttct taaagaaata gagacggggc cgcactatgt tgaccaggct ggtctcgaac 720  
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<210> 17  
<211> 1001  
<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 501

<223> 12-456-269 : polymorphic base A or G

<220>

<221> misc\_binding

<222> 481..500

<223> 12-456-269.mis1, potential

<220>

<221> misc\_binding

<222> 502..521

<223> 12-456-269.mis2, potential complement

<220>

<221> primer\_bind

<222> 233..252

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 693..712

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 12-456-269 potential probe

<220>

<221> misc\_feature

<222> 20,447

<223> n=a, g, c or t

<400> 17

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cagagccctg tagccgtggt aacctcaccc ctgtgctgga cgggaggcag gtcggctaag	180
cagaggtgct ggaagtgtgt gtggtaccag gactgggccg caggagctgc acagcctcac	240
agcacattag ccaatggccc catgccaaacc cccttggtcg ggttttctgg aacaggtacc	300
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aaaagatcac taagcaaag cctgtttgtc ctttgaata ctcaaccctg atcagtgggtg	420
aatactcagt caccaactat gtggaangag gcacaacccat ggagagacaac agatatggaa	480
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gcaattcatc tgggccaaat ccctcaggag cagggccatg cgggggttac aactgttcat	660
gcaacacaac agattgggct taagctggcc agggcagtga cgggcttttc atcttaattg	720
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<210> 18

<211> 1001

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

17

<222> 501  
 <223> 12-456-380 : polymorphic base G or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-456-380.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-456-380.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 122..141  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 582..601  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-456-380 potential probe

<220>  
 <221> misc\_feature  
 <222> 336  
 <223> n=a, g, c or t

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 aggtgatcca aaagatcact aagcaaatgc ctgtttgtcc tttggaatac tcaaccctga 300  
 tcagtgggtga atactcagtc accaactatg tggaangagg cacaaccatg gcagacaaca 360  
 gatattggaaa gagcaggaat tcttctagca ggaattctta tatcaaatgc aatggtggcc 420  
 atgcccccaa cagctctgca gttggaaaaa tgaagccttc aagtaaggcc aaagtcctac 480  
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<210> 19  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-457-204 : polymorphic base A or G

<220>

18

<221> misc\_binding  
 <222> 481..500  
 <223> 12-457-204.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-457-204.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 298..317  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 772..792  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-457-204 potential probe

<400> 19  
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<210> 20  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-457-206 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-457-206.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-457-206.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 296..315  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 770..790  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-457-206 potential probe

<400> 20  
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<210> 21  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-458-196 : polymorphic base T or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-458-196.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-458-196.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 679..696  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 200..217  
 <223> downstream amplification primer

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<220>
<221> misc_binding
<222> 489..513
<223> 12-458-196 potential probe

<220>
<221> misc_feature
<222> 33,792
<223> n=a, g, c or t

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<210> 22
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<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-458-438 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-458-438.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-458-438.mis2, potential complement

<220>
<221> primer_bind
<222> 921..938
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 442..459
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513

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<223> 12-458-438 potential probe

<220>
<221> misc_feature
<222> 275
<223> n=a, g, c or t

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atgaagtcac caaagcaaag attgcataac tgatgcatag gcctatcttg tgttatactg      180
ggagacaggc caatgtttcc attaatagac aagagcacca ccacgctgcc aaatggagct      240
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<220>
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<223> 12-460-274.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-460-274.mis2, potential complement

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<222> 760..777
<223> downstream amplification primer, complement

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<222> 489..513
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<220>
<221> misc_feature

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&lt;222&gt; 873

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 23

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&lt;210&gt; 24

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-461-124 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-461-124.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-461-124.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 378..396

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 911..928

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-461-124 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 850

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 24

23

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&lt;210&gt; 25

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-461-299 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-461-299.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-461-299.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 203..221

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 736..753

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-461-299 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 675,831,997

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 25

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24

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&lt;210&gt; 26

&lt;211&gt; 985

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-461-465 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-461-465.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-461-465.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 37..55

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 570..587

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-461-465 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 509,665,831

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 26

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25

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&lt;210&gt; 27

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-462-280 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-462-280.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-462-280.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 222..241

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 655..675

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-462-280 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 174

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 27

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26

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&lt;210&gt; 28

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-464-66 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-464-66.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-464-66.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 436..455

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 880..900

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-464-66 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 121,163..164

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 28

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1001

&lt;210&gt; 29

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-465-26 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-465-26.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-465-26.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 476..493

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 945..962

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-465-26 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 556,630,796

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 29

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&lt;210&gt; 30

&lt;211&gt; 1001

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<213> Homo Sapiens

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<221> allele
<222> 501
<223> 12-465-234 : polymorphic base G or T

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<222> 481..500
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<220>
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<222> 502..521
<223> 12-465-234.mis2, potential complement

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<222> 266..283
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 735..752
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-465-234 potential probe

<220>
<221> misc_feature
<222> 346,420,586,869..870,876..877
<223> n=a, g, c or t

<400> 30
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tcagagactc tctggacctg agagagaaca gtaactgtat tcaggcctaa tccctcagat      780
ggctgccttg gcggccactc tctttaccat gaaaattgcc cccagatgg aaacctcttg      840
aacttggaaa aacctcttta tttccaaann cagaannaaa aaaaaatcac tatgtatttc      900
ttcttaagaa aaaaaaattg ttattcaact actgtgtccc attttcccaa taaacattaa      960
ttgaactgct agtgggcacc attcatccat ttaacaaata c                                1001

<210> 31
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>

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29

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<221> allele
<222> 501
<223> 10-428-219 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 10-428-219.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-428-219.mis2, potential complement

<220>
<221> primer_bind
<222> 278..295
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 613..632
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-428-219 potential probe

<220>
<221> misc_feature
<222> 203,238,334,348,369,374..376,645,675
<223> n=a, g, c or t

<400> 31
actcttttat atcataaagt aaaactactt tacagatgag gaaactatct atttgatgta      60
cacaataatt tctctaaatt ctttcagcta gtcagtgggt tgttgagcta gaatctgagg      120
aatccagatc ccatcctttc attgattatg tcactaagat aagtcacaga atcttctttt      180
agggaagttt tcaagaatag tangtaagct gctatcttaa aatattgatg tcagtagncc      240
tggaattttt accacgtatg caagcatggg ataagacaac cacaggtttc tcttctgtac      300
cagttaacct gttttatgat tatttcaagt tgtngaataa aagcttgnaa aattggcagg      360
atctgtttng aggnnnttac tcaaggatat ttattacttg tttcaggaac agcatctgtt      420
gcagttgcag gtctccttgc agctcttcga ataaccaaga acaaactgtc tgatcaaaca      480
atactattcc aaggagctgg rgaggatatt gccttgtgta atacttatgt tctcctaaac      540
taacatataa ttagtggttaa ttgctcatta actcaagatt tgacgtattc tttcaaattt      600
ttagtatatt atgactgtag gctttatgga agattcccaa attanagtac tgagacatat      660
taattattca taggnataat ttagtacaga agtaatgaga tattttttcac agttttaaat      720
ttgtggtatt atcaaagtgt gccgtaggta tttgttatgc atatctaagg tgccatacag      780
gaaaactttc caatgtatgt gtatatatgc aagcttttat gtatttaagc tggtttatct      840
gatactggct tttagtatac tccagtgtta caaaaacaag tttttaaaat ttgactcaca      900
attcagtgtg tatgtatgta actacatatg tgtttttaag aaaaaacatt ttgaattagg      960
gatttgtagt atattattac tgttcacatg caaaaatccc                        1000

<210> 32
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-429-84 : polymorphic base C or T

```

30

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<220>
<221> misc_binding
<222> 481..500
<223> 10-429-84.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-429-84.mis2, potential complement

<220>
<221> primer_bind
<222> 418..436
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 823..842
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-429-84 potential probe

<220>
<221> misc_feature
<222> 222
<223> n=a, g, c or t

<400> 32
taattataaa atttaccaaa taccaaggaa gttgattcct ttaatcttgc aataaatctt 60
acagcacttt tggatatcaa attagaagag taagtagatg cttggaaaca cagatctttt 120
atttttcttc agtttcattc tgttgccata ttggtcaagt tcccatccaa atgaccaagc 180
ttctttcttc aaattcaatt agagtagctt gaacctaaaca antaaacaat acagttcaat 240
gagcaagggt gaccagttca ttattttgaa ttcagtattt tttcttggag caaatcttgg 300
ttgtattaac aggattaaat aacttgacaa tcaaggatat caccagttta tctgtggcca 360
aatagcaagc cacataatag tggtctctgt gtagtaaaaa gggtaaataa ttgttatttg 420
gcttaacctt aatgtgacaa aaatattttg ttcccactaa caactttaaa attttttctt ttcactctg 540
atgtaaccct taataagata ycatgtagaa caactttaaa attttttctt ttcactctg 540
aggctgccct agggattgca cacctgattg tgatggcctt ggaaaaagaa ggtttaccaa 600
aagagaaagc catcaaaaag atatggctgg ttgattcaaa aggattaata gttaaggtaa 660
gaatttgctt tttttaacca gaataaagat aactatgcc tcttggatgg ccatctcaaa 720
gacaactctg tttctaccac ctccctttac actatattaa aagagtagcc attaaataca 780
aatgatcatg tcatacagaa tactgatact cttctgaata aagaaaacta tgacttagga 840
acagattcca tctgcaatgt gagtcggctc cttacacccc tttgaattga cagtccattt 900
gaattagttt agttttcttt ctttctttcc ttcctttctt tcctttctct ttctttcttt 960
ctttttcttt ctttctttcc ttccttctct tccttctttt t 1001

<210> 33
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-420-284 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 10-420-284.mis1, potential

```

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-420-284.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 216..235  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 646..665  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-420-284 potential probe

<220>  
 <221> misc\_feature  
 <222> 42,308,413  
 <223> n=a, g, c or t

<400> 33  
 agcctatcgg ttgaaccact gcagagaata gagtgatggt cnttagggca tcctgtactt 60  
 tgcattgctcc tcctggaagt aaagagtaag acagagaata gtaataatca cccattccag 120  
 aactgggtgc acaacatcac aaaagcttgt ccagacttat tagcaagta ataaaaaact 180  
 agacttcctt ctaagtactt ataatttagg ctgtggggta gttctgttat gatacatttg 240  
 ttttaaaata ttctgcttct ttttaaagtg agttgtatgt gtctttgttg tagggacgtg 300  
 caattttntg ccagtggcag tccttttgat ccagtcactc ttccaaatgg acagacccta 360  
 tatectggcc aaggcaacaa ttcttatgtg ttccctggag ttgctcttgg tgnttggtggc 420  
 gtgtggattg aggcagatca cagataatat ttctctact actgctgagg tattgtaaaa 480  
 tcctctaagt ttaccaaggg yttaaaatac caagtgtgct cagcctaggt tgtctaattgt 540  
 tttatttatc tagcatctca gcttactctc tgaaagaagt aaagtctgaa gaacttccca 600  
 gtggagtata aggggtgggt agcatgttca tactgactca caaacgaaag gttcttcttc 660  
 agtagtcatt agaaaaattg tgtttttgat ttcttaagag gaacattttt gtgtcttcac 720  
 acatcagatc aagttctgtg acagtgtgag gacaattaaa aatattgttt ccagggtctgg 780  
 gtgtggtggc tcatgccct aatcccagca ctttgggagg ctgaggcagg cagatcactt 840  
 gagcccagga gttcgaggcc agcttgccca acatggtgaa acttcatctc tactaaaaat 900  
 acaaacatta gccgggcatg gtggtgtgtg cctgtgatcc cccctgactc aagaggctga 960  
 ggcaggagaa ttgcttgaat ccaggaggca gaggttgagc t 1001

<210> 34  
 <211> 821  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-423-411 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-423-411.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521

<223> 10-423-411.mis2, potential complement

<220>

<221> primer\_bind

<222> 91..109

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 510..528

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 10-423-411 potential probe

<220>

<221> misc\_feature

<222> 5,804

<223> n=a, g, c or t

<400> 34

cctangattc	tattcccatc	tttttcacca	attatgtgat	ctcaaccacc	accacaccac	60
tctgggctta	tttctttggt	gatagaagtg	gggatgaaat	taggtaatca	tattaatgtc	120
actgggtttt	gtatagacca	ttatatgtaa	ccgttcactt	agttattggc	gtaccaccca	180
cattgtatcg	tgtgaagtat	aaacactttt	tttctttctt	tttttcataa	ggttatagct	240
cagcaagtgt	cagataaaca	cttgggaagag	ggtcggcttt	atcctccttt	gaataccatt	300
agagatgttt	ctctgaaaat	tgcagaaaag	gtaaaaccac	tcttgttcaa	gcttcattat	360
ttttccttcc	ttttcttget	aaatatgcat	ttttaaatat	taaaaatctg	cttccttgaa	420
atgtatatct	gtatcaactt	actatgtcag	aagtcagaga	aatgaggcac	actgacactg	480
tagagcttag	gagctacatt	ygcttctcag	agaagaatgg	aagtattggg	gccgaataat	540
taatttcctt	ctctatcggt	ctttttttct	cctaaaattt	aaataattca	ttagggcttc	600
ctgtcagcca	gatttctccc	atttcacctt	tttaaagttg	ttcttttcta	tatcacatta	660
gtgataatta	accaaaatat	tacagatatg	ttagagtaac	atttagacct	tgaaccacct	720
tccatatcca	tgtcactatg	atgtgtggca	tcataaatga	cagatcacia	ctgataatac	780
tgggcggaag	gcaaaataaa	gagntcaaag	aacttgaaag	c		821

<210> 35

<211> 732

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 232

<223> 12-713-95 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 213..231

<223> 12-713-95.mis1

<220>

<221> misc\_binding

<222> 233..252

<223> 12-713-95.mis2, potential complement

<220>

<221> primer\_bind

<222> 137..153

<223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 586..604  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 220..244  
 <223> 12-713-95 potential probe

<220>  
 <221> misc\_feature  
 <222> 154  
 <223> n=a, g, c or t

<400> 35  
 ggtctcttct gtgaccccat tattgtcctg tgcatttatt tcaagatcag attcattatt 60  
 aaatctatcc agacaattcc tctatcatgt ccttgatcca tttctataac cataaccag 120  
 tttctataat cataacccaa cttcaccatt ttantatctt tacaccatct cggtaacttc 180  
 tgattttctt taggatcata ttaataagta ttcaggaact tacctttacc cygctattga 240  
 ctgttttcac ctacatataa atccctttgt gatctggccc atgcacttca cttcctgtc 300  
 tgccatcctc caccctccg tgtatccttt acaccattgt ataaataaat agttttagtt 360  
 ggctgcacat gtcattgttc tgtcacactt cttctctgta taaactgctc cttcttccc 420  
 cagtgtcttt gcttatttaa ttcattgcta tctttcataa ctcagggtta atatctagac 480  
 tggattagag aagttttctg tatagtatcc tagcatggat acaccaaagt agcatgtatc 540  
 aactgtatt gtaattatat gtatttattt gacttagcat taagtcacat agatattaga 600  
 ggtcagggcc tcattctttt ctctcttctt ttataatata tgttggtact gtgatacttc 660  
 tttttaataa ctttaatttt aagttcagga gtacatgtgc aggtttgtta cataggtgac 720  
 cttgtatgat gg 732

<210> 36  
 <211> 786  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 286  
 <223> 12-713-149 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 266..285  
 <223> 12-713-149.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 287..306  
 <223> 12-713-149.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 137..153  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 586..604  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 274..298

<223> 12-713-149 potential probe

<220>

<221> misc\_feature

<222> 154

<223> n=a, g, c or t

<400> 36

gggtctcttct	gtgaccccat	tattgtcctg	tgcatTTatc	tcaagatcag	attcattatt	60
aaatctatcc	agacaattcc	tctatcatgt	ccttgatcca	tttctataac	cataacccag	120
tttctataat	cataacccaa	cttcaccatt	ttantatctt	tacaccatct	cggtaacttc	180
tgatttttctt	taggatcata	ttaataagta	ttcaggaact	tacctttacc	ctgctattga	240
ctgttttcac	ctacatataa	atccctttgt	gatctggccc	atgcasttca	ccttcctgtc	300
tgccatcctc	cacccctccg	tgtatccttt	acaccattgt	ataaataaat	agtttttagtt	360
ggctgcacat	gtcattgttc	tgtcacactt	cttctctgta	taaactgctc	cttcttcccc	420
cagtgtcttt	gcttatttaa	ttcatgctta	tctttcataa	ctcagggttaa	atatctagac	480
tggattagag	aagttttctg	tatagtatcc	tagcatggat	acaccaaaaat	agcatgtatc	540
acactgtatt	gtaattatat	gtatttattt	gacttagcat	taagtcatca	agatattaga	600
ggtcagggcc	tcattctttt	ctctcttctt	ttataatata	tgttgttact	gtgatacttc	660
tttttaataa	ctttaatttt	aagttcagga	gtacatgtgc	aggtttggtta	cataggtgac	720
cttgtatgat	gggggtttgt	tgtacagatt	atttcatcac	ccaggaacta	agcatagtac	780
tcgtta						786

<210> 37

<211> 902

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 500

<223> 12-716-295 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 477..499

<223> 12-716-295.mis1

<220>

<221> misc\_binding

<222> 501..520

<223> 12-716-295.mis2, potential complement

<220>

<221> primer\_bind

<222> 206..225

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 727..746

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 488..512

<223> 12-716-295 potential probe

<220>

<221> misc\_feature

<222> 67,139,690,834

<223> n=a, g, c or t

35

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<400> 37
caatgtgtac tacaaaaata aaatgcatgc aatttagagt cactattgca ataattatga      60
actgtgntaa gtcatatatt atgaaaaagc cactcttcta tacattactt aagggtgaaaa      120
tctatgcaaa aaatatggnt agtgtatgta gagaatatat catgacatga ttagaatgat      180
atthtccccca tgcaagatat attcactaga gtcattaaac agatgaaaga gaattttaaac      240
tagaaagaga ggtatagata tccacttact aaataagaag ttattcagaa atagtagctc      300
aagctatctc tctcttgggt tcttttgaag taaactgccc agaagtttgg atctggggaa      360
ctctaagctc ccttccaact ctgagattct atgattttaa gggtttctat aagattttaa      420
cagttgaagc atcaaagttg aagtatcatc agcaatcact gtatttgtat tattattttt      480
atthacatcg aattggacay ttacggtcag atagacagaa aatattctaa aatattcctat      540
ttggtacctt aatttcatat gggttatgcta gatcacttaa aactcaaaat gtgttaagt      600
catacattat atttaaagat atatttcata atatttctcg agaacacaat gaaaacacta      660
atcttccatt tatagaatta attaataaan atcagaaaga tatttttagc actatcaaaa      720
atttaacttg aacatgattc ttgtacatgg cctggtaacta ccttgattta aatttcattg      780
cagagaaggg aatacatgta gaaaattgaa atttcataaa actgtaaatt ttnacaata      840
aattattcca taataagttt tactttaaga tttaaaacta agctttttat aagtcagaca      900
ga                                                                902

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<210> 38
<211> 981
<212> DNA
<213> Homo Sapiens

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```

<220>
<221> allele
<222> 480
<223> 12-720-80 : polymorphic base G or C

```

```

<220>
<221> misc_binding
<222> 461..479
<223> 12-720-80.mis1

```

```

<220>
<221> misc_binding
<222> 481..500
<223> 12-720-80.mis2, potential complement

```

```

<220>
<221> primer_bind
<222> 400..419
<223> upstream amplification primer

```

```

<220>
<221> primer_bind
<222> 856..876
<223> downstream amplification primer, complement

```

```

<220>
<221> misc_binding
<222> 468..492
<223> 12-720-80 potential probe

```

```

<220>
<221> misc_feature
<222> 191,218,225,437,480,694,772,775
<223> n=a, g, c or t

```

```

<400> 38
ctgcaggaaa catttggcaa tgtctggaag ctttgagttg tcttaactag ggtagcgggg      60
ttggaagggg gtaaaggag gggctactga catctagcag atagaggcag cggaatgatt      120
ctactaaata tctgtagt cactggacag cccctgcgac aaggaattat ccagcctaaa      180
atgtcaatag ntgctgaggt tgagaaatac tgcagtanga taagnaccct cctggaagac      240

```

36

agggatcaca tctcacaat ctttattttc cttatagcac ttaatacagt acttacacaa	300
tgcattgtgca gtgcaagact ttaaagtggg tttagaaaac tcaaaggaat ttatgaaact	360
ttattcttta ttttgtgcat atataatttt gattttttacc ttgtattata ctccattcct	420
acaagctctt gtgaaancag atgacaacgt tgaaaataat gatgataact acaataaaas	480
gaagtactta ctccatgaca ggaactttac taagtgttta atattgtttt tacaaaatat	540
attctgtcaa taatcttatt gttttttcatt tccattatgc taattgcttt attcttctgt	600
taaagttact tcagtgggtg gtactttact tttgcagatg aatcagatgg ttgtcttaag	660
tgatagcgtt gggtttataaa acttaattgc tttngtagcc tgacaagact gaaatatact	720
catagaattg cttataaaga aagttttacac attagataca aatttccaag tnganttgg	780
aactaactca atgtactatg gaagccaaat attatcttta gtttttaata gatatttgta	840
gatttgtagc cctaaggatt gtaaggattg ttggcattaa caaacttgcc tctttttgtc	900
actatacttg aatacaaata gtttatatag cattttgttt aggggtctatt agaaaatgag	960
ctgatagata tcattgattt g	981

&lt;210&gt; 39

&lt;211&gt; 872

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 372

&lt;223&gt; 12-721-227 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 352..371

&lt;223&gt; 12-721-227.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 373..392

&lt;223&gt; 12-721-227.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 146..165

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 588..607

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 360..384

&lt;223&gt; 12-721-227 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 756,794,799

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 39

aaattaatga gttaaatgat gagtttatta taaaattatg gtcaaacacc tacaattata	60
tgttataata ttttgtataa tatgtagagc aatactaadc aaacaaatag ttatactagt	120
aactttacct ccttctcctt tccatcttac ctctgcctaa atgtccctgg tcagaagtcc	180
ttagtcaaccg acgggattta actccaaact ccttttttga cattgataat gctccagtcc	240
ttatttctgt acccagaccc tctgtctccc caggtcagga cacttggggc tgctgagtcc	300
agctcacaca caactctgcc tttggtctga ctctgttttt tgtctgaaat gcctacctac	360
tttctgtttg tytaactcca ggctttcctg taggtttgcc tttcaagcca ctgtgcacta	420
taaggccttc cagaactgct ctctctctgg atttcttagc acccacattt caccatttag	480



37

ctaagccgctc	ttctctttca	gcgttctctt	tttcttcccg	tactctgtta	ccttactaaa	540
ttatacatct	ctcaaggaca	ataacacttt	tcttaagctt	ctccgtagtg	ttatgcttgt	600
tcagtgcata	tatgtttgct	gatggcaaag	aataatat	tggaaataat	ttcatgattt	660
aaaaatctaa	aagatattaa	gatatactga	aaaataagta	tttgattatt	taaaatgtta	720
caaaagagga	aggtttccta	ctccctctct	gcatcncaca	ttttcatgga	acaaaggcct	780
aggataaagc	tgantttang	atttgccccc	atctagaaat	ttaatcaaag	tcttagagct	840
ggagagaaca	tcagggttac	tgagtctccc	tc			872

&lt;210&gt; 40

&lt;211&gt; 926

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 426

&lt;223&gt; 12-721-281 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 406..425

&lt;223&gt; 12-721-281.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 427..446

&lt;223&gt; 12-721-281.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 146..165

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 588..607

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 414..438

&lt;223&gt; 12-721-281 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 756,794,799,893

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 40

aaattaatga	gttaaatgat	gagtttatta	taaaattatg	gtcaacaccc	tacaattata	60
tggtataata	ttttgtataa	tatgtagagc	aataactaatc	aaacaaatag	ttatactagt	120
aactttacct	ccttctcctt	tccatcttac	ctctgcctaa	atgtccctgg	tcagaagtcc	180
ttagtcaccg	acgggattta	actccaaact	ccttttttga	cattgataat	gctccagtcc	240
ttatttctgt	accagagacc	tctgtctccc	caggtcagga	cacttgggcc	tgctgagtcc	300
agctcacaca	caactctgcc	tttggctctga	ctctgttttt	tgtctgaaat	gcctacctac	360
tttctgtttg	tctaactcca	ggctttcctg	taggtttgcc	tttcaagcca	ctgtgcacta	420
taaggmcttc	cagaactgct	cttctctctg	atttcttagc	acccacattt	caccattagc	480
ctaagccgctc	ttctctttca	gcgttctctt	tttcttcccg	tactctgtta	ccttactaaa	540
ttatacatct	ctcaaggaca	ataacacttt	tcttaagctt	ctccgtagtg	ttatgcttgt	600
tcagtgcata	tatgtttgct	gatggcaaag	aataatat	tggaaataat	ttcatgattt	660
aaaaatctaa	aagatattaa	gatatactga	aaaataagta	tttgattatt	taaaatgtta	720
caaaaagagga	aggtttccta	ctccctctct	gcatcncaca	ttttcatgga	acaaaggcct	780
aggataaagc	tgantttang	atttgccccc	atctagaaat	ttaatcaaag	tcttagagct	840

38

ggagagaaca tcagggttac tgagtctccc tctctgcctt agggttcaat acnaaaattt 900  
 aacactgttt gatttggaac taggga 926

<210> 41  
 <211> 1000  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-721-440 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-721-440.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-721-440.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 62..81  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 504..523  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-721-440 potential probe

<220>  
 <221> misc\_feature  
 <222> 672,710,715,809  
 <223> n=a, g, c or t

<400> 41  
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 caaactcctt ttttgacatt gataatgctc cagtccttat ttctgtaccc agaccctctg 180  
 ctccccagg tcaggacact tgggcctgct gaggccagct cacacacaac tctgcctttg 240  
 gtctgactct gttttttgtc tgaaatgcct acctacttct tgtttgctca actccaggct 300  
 ttctgttagg tttgccttct aagccactgt gcactataag gccttcacaga actgctcttc 360  
 ctctggattt cttagcacc acatttcacc attagcctaa gccgtcttct ctttcagcgt 420  
 tctcttttct tctccgtact ctgttacctt actaaattat acatctctca aggacaataa 480  
 cacttttctt aagcttctcc rtagtgttat gcttggtcag tgcataatag tttgctgatg 540  
 gcaaagaata atattttgga aataatttca tgatttaaaa atctaaaaga tattaagata 600  
 tactgaaaaa taagtatttg attatttaaa atgttacaaa agagggaagg ttctactctc 660  
 ctctctgcat cncacatttt catggaacaa aggcctaggg ataagctgan tttangattt 720  
 gcccccatct agaaatttaa tcaaagtctt agagctggag agaacatcag gggtactgag 780  
 tctccctctc tgccttaggg ttcaatacna aaatttaaca ctgtttgatt tgggaactagg 840  
 gaatggtttg gggacagtaa atgttgaggc taattagggtg atttagataa tccagagtct 900  
 gtgtaaaatt aaaatcctat ttgtagtgag actctgaact agctttctca gccttttgct 960  
 ctacctctg cacaagaata tgactcagag ctgggagtaa 1000

<210> 42

39

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<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 12-723-293 : polymorphic base C or T

<220>
<221> misc_binding
<222> 484..502
<223> 12-723-293.mis1

<220>
<221> misc_binding
<222> 504..523
<223> 12-723-293.mis2, potential complement

<220>
<221> primer_bind
<222> 210..230
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 591..610
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 12-723-293 potential probe

<220>
<221> misc_feature
<222> 312,379
<223> n=a, g, c or t

<400> 42
ctcattagtt ctcagacatt tgaatagtc ttctccagtg catagttact tgggtaaagg      60
gctgtagttc cctgtggata aagattaggg ggattattac cctgtattct tgtcatgggt      120
ttataagggt ctttcttctg ggaactaggc ctcttcttca gtttatgggt tctgggtcta      180
aaaattaact caggttcgga agctgagcaa gagattgtga cttcattggg gcagctacac      240
tcaactttgt ggtcattcat ctttggtttc ttttgatagt aaatattgag tagtagcttt      300
gttagctctc cngttttgcc tctagggatg caagattccg ttaattatct caaaaacatt      360
ttgtgggtta gccacacanc accccacttt tgccactctt atgataattg tgtccatctg      420
gcttctgatt gttaagcatt ctcacctggg cttagcatcc ttattgctat cagtcaggct      480
ggttatatat catccttttc ttytgctact cctgaccttc aggggagaa caccactgaa      540
attttttagta atgttgatgt cttctcattg gcatattcct tatggccttg ggaaatgggt      600
tgttttctgg gccttcctgt ggaatatgtt catcttatgg attttctggc ttttacatgt      660
gtctatatac cagactgccc agttctctga gcctcgcaat cccctcttct gtcatttgcc      720
acagcattcc tggcatttca gtcataatag tatcagccat taatttgtcc atgtttctag      780
gaaccatcct agcaatgagt ttgcaccatc ttctgggatt cttgccacat gttaaatcct      840
gtatctcagg agagtgcctg caaatcagta agctctccct tttcctagggt tccagccctt      900
ttaatcaaac accctcagaa ttttaattcta catgtattct cctggctaata acaaatactg      960
gtcttgtggc tcctttgggg taaagtctct ttcttccttt a                                1001

<210> 43
<211> 1001
<212> DNA
<213> Homo Sapiens

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<220>
<221> allele
<222> 501
<223> 12-724-195 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-724-195.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-724-195.mis2, potential complement

<220>
<221> primer_bind
<222> 307..326
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 797..817
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-724-195 potential probe

<220>
<221> misc_feature
<222> 244,249,631,669
<223> n=a, g, c or t

<400> 43
tcacctgagg tcaggagttc gagaccagcc tggccaacat ggtgaaaccc tgtctctact      60
aaaaatacaa aaattagcta ggcgtgggtg caggcgcttg taatcccagc tacttggggag      120
gctgaggcag gagaatcact tgaacccagg tgggtggagg tgcaagtgag cgtgattgca      180
ccaccgcact ccagcctggg caacagagcg agaatccatc tcaaaaaaaaa aaaaaaaaaag      240
aaanaagana aagcatgaag tatgtggagg catacatgaa aagagacaaa tgacatgtta      300
tttttacatt ttaataggca accaagtata acaataatgt gaagcaagac tttctaatta      360
tgtgcacagg ccatctgatg gacggcaaaag gtggaacatg gggagatgag tatgtgaaaag      420
agggaaacatc atggcctctg agctgaaggc agcatgggta cagggtcaaa tcctcagaga      480
agctttggct ctaaaggact yatctgtgct tcaacatacc actgtattct cctatttcta      540
gtttgtacca ttccttatgc ctaaaatgcc ttcatataag ggatgcagaa atttagaaaa      600
tgcttttata aactccatgt gcaaacttac naatagagaa ttaaagggtct cctcaaagca      660
tgttccagnt aaatacctca tctccaacct caatcatgtc atttacttct gtagtttgaa      720
tcactaaatc tgtgactttt tctccattta gcctgaatca aactgtttat ttctaaaagc      780
cagcaacttc tatgaacact aagggtcaca tattcagcag aaaagagaat tgtatattat      840
tttactgaag aatattacag tcataaaatt ttagatctag aaatgtcctc agagatgatt      900
tagcaataac taatgaccag gaagggttaa aaacacccaa ggttcagaa aaatatatgg      960
tagtgagtgt tgccatctaa gagaaaagcc agaggggacta a                               1001

<210> 44
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-724-225 : polymorphic base C or T

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<220>
<221> misc_binding
<222> 482..500
<223> 12-724-225.mis1

<220>
<221> misc_binding
<222> 502..521
<223> 12-724-225.mis2, potential complement

<220>
<221> primer_bind
<222> 277..296
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 767..787
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-724-225 potential probe

<220>
<221> misc_feature
<222> 214,219,601,639
<223> n=a, g, c or t

<400> 44
tggccaacat ggtgaaaccc tgtctctact aaaaatacaa aaattagcta ggcgtggtgg      60
caggcgcttg taatcccagc tacttgggag gctgaggcag gagaatcact tgaacccagg      120
tggtggaggt tgcagtgagc cgtgattgca ccaccgcact ccagcctggg caacagagcg      180
agaatccatc tcaaaaaaaaa aaaaaaaaaaag aaanaagana aagcatgaag tatgtggagg      240
catacatgaa aagagacaaa tgacatgtta tttttacatt ttaataggca accaagtata      300
acaataatgt gaagcaagac tttctaatta tgtgcacagg ccatctgatg gacggcaaag      360
gtggaacatg gggagatgag tatgtgaaag agggaaacatc atggcctctg agctgaaggc      420
agcatgggta caggctcaaa tcctcagaga agctttgggt cttaaaggact tatctgtgct      480
tcaacatacc actgtattct yctatttcta gtttgtacca ttccttatgc ctaaaatgcc      540
ttcatataag ggatgcagaa atttagaaaa tgcttttata aactccatgt gcaaaacttac      600
naatagagaa ttaaagggtct cctcaaagca tgttccagnt aaatacctca tctccaacct      660
caatcatgtc atttacttct gtagtttgaa tcaataaatc tgtgactttt tctccattta      720
gcctgaatca aactgtttat ttctaaaagc cagcaacttc tatgaacact aagggtcaca      780
tattcagcag aaaagagaat tgtatattat ttactgaag aatattacag tcataaaatt      840
ttagatctag aaatgtcctc agagatgatt tagcaataac taatgaccag gaagggttaa      900
aaacacccaa gggtccagaa aaatatatgg tagtgagtgt tgccatctaa gagaaaagcc      960
agagggacta agtccaatag ccacgtacgg tagagcagac c                                1001

<210> 45
<211> 421
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 330
<223> 10-153-329 : polymorphic base G or T

<220>
<221> misc_binding
<222> 310..329

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<223> 10-153-329.mis1, potential

<220>
<221> misc_binding
<222> 331..349
<223> 10-153-329.mis2, complement

<220>
<221> primer_bind
<222> 1..20
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 402..421
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 318..342
<223> 10-153-329 potential probe

<400> 45
agagtcaccc tbggctcttag gtagtaggtg gagctgaggg ataatggccc aaggccaaga      60
gttgatcctt ccaactttgt tcagtgatcc agctttcata tcaggatgatc aggacaacca      120
ggccaatctg atagggggcg gtgtttataa aaaggccact cacctagagc cagaagctcc      180
acaccagcca ttacaaccct gccaatctca agcacctgcc tctacaggta cctttcttgg      240
gaccaattta caatctcttg gatccccaac tatagaacct ggaagctagt ggggacagaa      300
agacggggag cctgggctag gtgtaggggk cctgagttcc gggctttgct acccagctct      360
tgacttctgt ttcccgattt taaatgagca gtttggaacta agccattttt aaggagagcg      420
a                                                                421

<210> 46
<211> 428
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 342
<223> 10-95-342 : polymorphic base A or G

<220>
<221> misc_binding
<222> 322..341
<223> 10-95-342.mis1, potential

<220>
<221> misc_binding
<222> 343..361
<223> 10-95-342.mis2, complement

<220>
<221> primer_bind
<222> 3..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 404..422
<223> downstream amplification primer, complement

<220>

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<221> misc_binding
<222> 330..354
<223> 10-95-342 potential probe

<220>
<221> misc_feature
<222> 192,253..254,424..425
<223> n=a, g, c or t

<400> 46
rwtccctcct tttttccctg cagttggtac agatggcatt gtcccagtct gttcccttct      60
cggccacaga gcttctcctg gcctctgcca tcttctgcct ggtattcttg gtgctcaagg      120
gtttgaggcc tcgggtcccc aaaggcctga aaagtccacc agagccatgg ggctggccct      180
tgctcgggca tntgctgacc ctggggaaga acccgcacct ggcactgtca aggatgagcc      240
agcgctacgg ggnngtcctg cagatccgca ttggctccac gcccgctgctg gtgctgagcc      300
gcctggacac catccggcag gccctggtgc ggcagggcga cratttcaag ggccggcctg      360
acctctacac ctccaccctc atcactgatg gccagagctt gaccttcagc acagactctg      420
gagnnata                                     428

<210> 47
<211> 373
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 277
<223> 10-100-277 : polymorphic base C or T

<220>
<221> misc_binding
<222> 258..276
<223> 10-100-277.mis1

<220>
<221> misc_binding
<222> 278..297
<223> 10-100-277.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 355..372
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 265..289
<223> 10-100-277 potential probe

<400> 47
tagggatgga gatggcggtg ggcaggctgt ctggatgggg tggaggtagg agcaacacat      60
gccccagctt tccagccctg agcctcacag tgccctcttc cctcctcagc acaacaaggg      120
acacaacgct gaatggcttc tacatcccca agaaatgctg tgtcttcgta aaccagtggc      180
aggtaacca tgaccctgta gtacataccc ctcacgaaaa aatgtgtgca ggttcagcag      240
tcaggaaggc tgtttgctcc tgctaggaac tgtttayata atgaaaggag gggacctcaa      300
ttgctatagt ctgctctaag tgacgatatt tacaaaagtt tcacaaactt tagtgcacag      360
gaatcaacta ggg                                     373

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<210> 48  
 <211> 375  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 297  
 <223> 10-102-294 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 278..296  
 <223> 10-102-294.mis1

<220>  
 <221> misc\_binding  
 <222> 298..317  
 <223> 10-102-294.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 356..375  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 285..309  
 <223> 10-102-294 potential probe

<220>  
 <221> misc\_feature  
 <222> 30,218  
 <223> n=a, g, c or t

<400> 48	
agagwgctgt gggaggaccc ctctgagttt cggcctgagc gggttcctcac cgccgatggc	60
actgccatta acaagccctt gactgagaag atgatgctgt ttggcatggg caagcgccgg	120
tgtatcgagg aagtccctggc caagtgggag atcttcctct tcctggccat cctgctacag	180
caactggagt tcagcgtgcc gccgggcgtg aaagtcgncc tgaccccat ctacgggctg	240
accatgaagc acgcccgtg tgaacatgtc caggcgccgc tgcgcttctc catcaaytga	300
agaagacacc accattctga ggccagggag cgagtggggg ccagccacgg ggactcagcc	360
cttgtttctc ttcct	375

<210> 49  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-413-394 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 10-413-394.mis1



<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 10-413-394.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 108..125  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 539..556  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-413-394 potential probe

<400> 49  
 tgtgatttgt gtaccaattg cctgggtcat tgcgtggcac atcacaggcc atctataagt 60  
 ggcagctata acaatcacca tcacatttat gtacaaaatt cagaaatatac gaatctatgt 120  
 gtggcaataa tgaacattaa aaaatacaat gaaaatgtca gtctgaatca tacatagtat 180  
 ttggagcaaa tagcgactta ttttgcctgt atttgcattt cctttcccag ttctcaaaaag 240  
 tctatggtcc tgtgttcacc gtgtattttg gcatgaatcc catagtgggtg tttcatggat 300  
 atgaggcagt gaaggaagcc ctgattgata atggagagga gttttctgga agaggcaatt 360  
 cccaatatc tcaaagaatt actaaaggac ttggtaggtg cacatatattc tgtgtcagct 420  
 ttggttaactg ggggtgagggg gatggaaaac agagccctaa aaagcttctc agcagagctt 480  
 agcctatctg catggctgcc ragtggtgca gcactttctt ccttggctgt gaattctccc 540  
 agtttctgcc ccttttttta ttaggaatca tttccagcaa tggaaagaga tggaaggaga 600  
 tccggcgttt ctccctcaca accttgccgga attttgggat ggggaagagg agcattgagg 660  
 accgtgttca agaggaagct cactgccttg tggaggagtt gagaaaaacc aagggtgggt 720  
 gactctactc tgcgtcattg accttaacag ttacctgtct tcactagtga cgtccttgga 780  
 aacatttcag ggggtggccag gtcttcattg cgcctcctgg ttgtcagccc tcagggtggtg 840  
 gagggagatt tgaagcacag agacaaggga ggttttgtgt atctgtgctt tgccctgtata 900  
 aatgtgttgg ttcataagggt gtaggaataa aaggctcattt aatcctattt tctgctcaat 960  
 tttgtttcct tgttttcaaa tcagaaatta taaataaggg t 1001

<210> 50  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-414-243 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-414-243.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-414-243.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 259..276

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<223> upstream amplification primer

<220>
<221> primer_bind
<222> 592..609
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-414-243 potential probe

<220>
<221> misc_feature
<222> 981
<223> n=a, g, c or t

<400> 50
atgaaaatgt cagtctgaat catacatagt atttggagca aatagcgact tattttgctg      60
ctatttgcac ttcctttccc agttctcaaa agtctatggc cctgtgttca ccgtgtattt      120
tggcatgaat cccatagtgg tgtttcatgg atatgaggca gtgaaggaag ccctgattga      180
taatggagag gagttttctg gaagaggcaa tcccccaata tctcaaagaa ttactaaagg      240
acttggtagg tgcacatatt tctgtgtcag ctttggtaac tggggtgagg gggatggaaa      300
acagagccct aaaaagcttc tcagcagagc ttagcctatc tgcattggctg ccaagtgttg      360
cagcactttc ttccttggct gtgaattctc ccagtttctg cccctttttt tattaggaat      420
catttccagc aatggaaaaga gatggaagga gatccggcgt ttctccctca caaccttgcg      480
gaattttggg atggggaaga rgagcattga ggaccgtgtt caagaggaag ctcactgcct      540
tgtggaggag ttgagaaaaa ccaagggtgg gtgactctac tctgcgtcat tgaccttaac      600
agttacctgt cttcactagt gacgtccttg gaaacatttc aggggtggcc aggtcttcat      660
tgcgcatcct gggtgtcagc cctcaggtgg tggagggaga tttgaagcac agagacaagg      720
gaggttttgt gtatctgtgc tttgcctgta taaatgtgtt ggttcatagg gtgtaggaat      780
aaaaggcat ttaatcctat tttctgtcga attttggttc cttgttttca aatcagaaat      840
tataaataag ggtcttgagt tcatttttga agagttaaag aaggtttccc ccatagcata      900
aatctgcatt acctccacca gaacatttca ccagagaaca cttggaaaag ggacatggtg      960
aaagtgcact catcacactt natgaaatat gaaccaagtt a                               1001

<210> 51
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-416-273 : polymorphic base A or T

<220>
<221> misc_binding
<222> 482..500
<223> 10-416-273.mis1

<220>
<221> misc_binding
<222> 502..520
<223> 10-416-273.mis2, complement

<220>
<221> primer_bind
<222> 229..246
<223> upstream amplification primer

<220>
<221> primer_bind

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<222> 630..649

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 10-416-273 potential probe

<220>

<221> misc\_feature

<222> 619,997

<223> n=a, g, c or t

<400> 51

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gacacaggag cccctgcatg caggatagga gccacatgcc ttacactgat gctgtagtgc	360
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gctataaaga ttataaang ctaggtctcc ttaataggct gctcttaggt gttacacttt	660
caaataattg ttgaaaatat cagtgtgtca atttcccaa acactcttct gagattttta	720
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gtaatgtctt attttcagtg cagtggttta caactctttt attagggttt tctagataa	900
ttaatatttt gttgtgttat aattttttaa ttttttttta tttctttgta attggtataa	960
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<210> 52

<211> 1001

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 501

<223> 10-418-177 : polymorphic base A or G

<220>

<221> misc\_binding

<222> 481..500

<223> 10-418-177.mis1, potential

<220>

<221> misc\_binding

<222> 502..520

<223> 10-418-177.mis2, complement

<220>

<221> primer\_bind

<222> 325..342

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 659..676

<223> downstream amplification primer, complement

<220>

<221> misc\_binding  
 <222> 489..513  
 <223> 10-418-177 potential probe

<220>  
 <221> misc\_feature  
 <222> 68,174,203,619,885,890,897,907,918  
 <223> n=a, g, c or t

<400> 52  
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 ctacataatc aaaatacaag atgtgtcaaa tttgaagtga tgaaatagag cggncaaatg 180  
 aggccagaaa agggcatcca aancttgatg atctggagaa cacattcaga aggttgacac 240  
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 ggggcttgca ggagtttaact ctgttggttc caattgagaa aatgaacatc ttggtttgac 840  
 tgaactctgc cactagatac atcactaagg cacccaagag ctccntcctn agaaggntaa 900  
 aattcantct gatctttnc ttaacctgct tgacaaatgt atctaagtcc acaactgcat 960  
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<210> 53  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-665-315 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-665-315.mis1, potential complement

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-665-315.mis2

<220>  
 <221> primer\_bind  
 <222> 795..815  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 357..377  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-665-315 potential probe

<220>  
 <221> misc\_feature  
 <222> 175,181..182,195,215,227,238,245,652,747  
 <223> n=a, g, c or t

<400> 53  
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 ttgaccttct ccccaccagc ctgccccatg cagtgtacgtg tgacattaaa ttcangaaac 180  
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 cataactcat atgtatggca gtttaactgg actttctctt gtttccagtt tggggctata 360  
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 tcaagattgg tttggctact cttagggtgt tatatttcca aataattttt aaagggtatta 480  
 gtttgtcaat ttcccaaaac yttgggctgg aatttctggc aggggtgacac taaatttata 540  
 ggctagtttg gaaagaactg aatcttgaca cgttgaggct ttccattcct gaataataatt 600  
 atgcttccaa tttgtttggg gtttctttta ttaaccagg aatgttgtga antttgttgt 660  
 catggctttc gagtcttttg tttccctag ataattaata tttttgttgt agaacataaa 720  
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<210> 54  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-666-324 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-666-324.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-666-324.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 186..205  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 621..641  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-666-324 potential probe

<400> 54  
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50

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atatacattt ctatgaatat cgtcccattg gctttaaaag agagtaaaca aattcacacg 240
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gagccaatat cttctaaaca gtgtttcagg tgtctggaat tcattagtga gccaaataaa 360
gttctctgct catataaaac atgtatttta gggatgaaac ataaacaata aaaataaatg 420
cagagaatta gcaaagttgt tagccacttc tactttccag tcagattgcc cttgttcaac 480
tgttggtctt gacacttaaa magctgtgtg accttgaaca agatacttaa cagatcatca 540
gtgcaattgt ttggttggtc aaattggtat aataataata tggtaggcta aataatgtct 600
tgacataaag agattaagct ggtgggtttc atgtaatcac aagggtcttt acattagtgt 660
taaatatagt gaaccccaag tttatcttca aagaatcagt atgtcagtat gtgcatctat 720
cttattgttt gattctccat tttaaagttt aacttcttaa ttctctttgc ccccttgctt 780
ccagtttcag taaacaactt tcttaccagt cctaataaat agttcacatc tgttccctg 840
gtcacctgct ttgacctag tccacctgg tcacctgctc tatcctgact catcctgagc 900
caccagttct gtaaccgccc tcccaccaa actacttacc ccaccactct ggctcatact 960
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&lt;210&gt; 55

&lt;211&gt; 436

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 155

&lt;223&gt; 10-72-155 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 135..154

&lt;223&gt; 10-72-155.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 156..175

&lt;223&gt; 10-72-155.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..20

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 417..436

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 143..167

&lt;223&gt; 10-72-155 potential probe

&lt;400&gt; 55

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tggacaaaat agtaacttcr tttgctgtta wctctrtcta ctttcctagc tctcaaargt 120
ctatggccct gtgttcaactc tgtatttttg cctgraaccc atrgtgggtgc tgcattggata 180
tgaaghrgtg aaggaagccc tgattgatct tggagaggag tttcttgga gaggcawttt 240
ccactggct gaaagagcta acagaggatt tggtaggtgt gcawgtgcct gtttcagcat 300
ctgtcttggg gatggggagg atggaaaaca gagacttdca gagctcctcg ggcagagctt 360
ggcccatcca catggctgcc cagtgtcagc ttcctcttctc ttgcctgggt ctcctccta 420
gtttcgtttc tcttcc

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436

&lt;210&gt; 56

&lt;211&gt; 678

&lt;212&gt; DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 178

<223> 10-76-177 : polymorphic base A or T

<220>

<221> misc\_binding

<222> 158..177

<223> 10-76-177.mis1, potential

<220>

<221> misc\_binding

<222> 179..198

<223> 10-76-177.mis2, potential complement

<220>

<221> primer\_bind

<222> 1..18

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 416..435

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 166..190

<223> 10-76-177 potential probe

<220>

<221> misc\_feature

<222> 549,635,642,655,661

<223> n=a, g, c or t

<400> 56	
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taacctgaaa tctctggttg acccaaagaa ccttgacacc actccagttg tcaatggwtt	180
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ggcatttctt tttctgcatg ttctaaataa aaagcattat tatttgctga gtcagtttat	600
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<210> 57

<211> 718

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 218

<223> 10-76-217 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 198..217  
 <223> 10-76-217.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 219..238  
 <223> 10-76-217.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 416..435  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 206..230  
 <223> 10-76-217 potential probe

<220>  
 <221> misc\_feature  
 <222> 549,635,642,655,661,708  
 <223> n=a, g, c or t

<400> 57  
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 taacctgaaa tctctggttg acccaaagaa ccttgacacc actccagttg tcaatggatt 180  
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 tgaacattcg acctccatta cggagagttt cctatgtttc actgtgcaaa tatatctgct 420  
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 ggcatttctt tttctgcatg ttctaaataa aaagcattat tatttgctga gtcagtttat 600  
 tagaccttcc ttcttttatg cataatgtag gtcangaaat tnaaagaaaa tagangttcc 660  
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<210> 58  
 <211> 834  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 334  
 <223> 10-76-333 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 315..333  
 <223> 10-76-333.mis1

<220>  
 <221> misc\_binding  
 <222> 335..354  
 <223> 10-76-333.mis2, potential complement

<220>



<221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 416..435  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 322..346  
 <223> 10-76-333 potential probe

<220>  
 <221> misc\_feature  
 <222> 549,635,642,655,661,708,751  
 <223> n=a, g, c or t

<400> 58  
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 agggaagcta ttttgggtga gtgttagagt ntacttgagg attggatttg aaagtgagaa 780  
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<210> 59  
 <211> 995  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-77-316 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-77-316.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-77-316.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 185..203  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 593..610

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 10-77-316 potential probe

<220>

<221> misc\_feature

<222> 427,513,520,533,539,586,629,762..765,774..776,813

<223> n=a, g, c or t

<400> 59

acctgaaatc tctgggtgac ccaaagaacc ttgacaccac tccagttgtc aatggatttg	60
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tggctgctgc tgtgcagtc ctgcagctct ctttcctctg gggcattatc catctttcac	180
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gagttattaa tatgttatta ttaaataagag aaatatgatt tgtgtattat aattcaaagg	420
catttctnttt tctgcatggt cttaaataaaa agcattatta tttgctgagt cagtttatta	480
gaccttcctt cttttatgca waatgtaggt cangaaattn aaagaaaata gangttccna	540
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ggaagctatt ttgggtgagt gttagagtnt acttgaggat tggatttgaa agtgagaaac	660
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cgccctccag gttcaagtga ttctcctgcc tcagcctcct gagtagctgg gattacaggt	900
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<210> 60

<211> 442

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 78

<223> 10-155-78 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 58..77

<223> 10-155-78.mis1, potential

<220>

<221> misc\_binding

<222> 79..98

<223> 10-155-78.mis2, potential complement

<220>

<221> primer\_bind

<222> 1..18

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 424..442

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

&lt;222&gt; 66..90

&lt;223&gt; 10-155-78 potential probe

&lt;400&gt; 60

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acttacctaa	gtactaaatg	ttataaaacc	aaactcttct	gacctctcaa	tctagtcaac	180
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gcactagcag	acagcatggt	cttggctaag	atactgaatc	ttcaaggctc	agcttcctca	420
ttccggaaat	gggtcaattt	ta				442

&lt;210&gt; 61

&lt;211&gt; 442

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 104

&lt;223&gt; 10-155-104 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 84..103

&lt;223&gt; 10-155-104.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 105..124

&lt;223&gt; 10-155-104.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 424..442

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 92..116

&lt;223&gt; 10-155-104 potential probe

&lt;400&gt; 61

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agtcagggac	caagtta	cttttctttg	ccctgtataa	aggsttctcc	aaggcctttg	120
acttacctaa	gtactaaatg	ttataaaacc	aaactcttct	gacctctcaa	tctagtcaac	180
tggggctgta	attattaatg	aaattaatgt	ttattttgaa	aataatttac	tagactgaat	240
tacgaaatcc	tgaatcattg	tacactatca	gtaaatattg	gtggacccaa	ctgaactgaa	300
tgttttgctt	gaaatgaaac	ctttgagatg	cagggcttat	gggttctagt	cccagctcta	360
gcactagcag	acagcatggt	cttggctaag	atactgaatc	ttcaaggctc	agcttcctca	420
ttccggaaat	gggtcaattt	ta				442

&lt;210&gt; 62

&lt;211&gt; 772

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

<220>  
 <221> allele  
 <222> 52  
 <223> 10-156-52 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 32..51  
 <223> 10-156-52.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 53..71  
 <223> 10-156-52.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 1..19  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 401..420  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 40..64  
 <223> 10-156-52 potential probe

<400> 62  
 ttcaaggctc agcytcctca ttccggaaat ggggtcaattt tattgtaagc araggtaatt 60  
 gagagattca aaagggacat gaggtgtaac aattctctgt aaattgtag aatccctgtt 120  
 aaaaatgacc agtaaagctt tgtgcaactg tgtcttgaca taactttatt tttcttaata 180  
 aaagaaatgg aaataacctc actagggaat ttagaacaaa tatgatgata tctttaaaga 240  
 aaatggcttt gcacaagtat tgacattaat gatctagtaa agtgatatctt tctagttgta 300  
 tttagatcct caactcagta tgtcagctcc tgtaaggctc tatacattgt ggtgggtctg 360  
 tgctgtgggt ccatttagtg atttccctac ctcccatctt ctattgcac cacaactgtg 420  
 gttctgtcca taatttcctt tgctttctgt gcattattac atcatatctg aaaatgagaa 480  
 accaaaaaca atagaaaagca gccatgtctg gaggtgactg ggggggtcgag aagccctagt 540  
 ttctcaaacc cttagcacca aatttttccc tcagttacac tgagcgtttc acttctgcag 600  
 tgatggagaa ggggatccc ttatttcttc tcattgagcat ctctgggtgct gtttccctta 660  
 gagacaaata aggggttcta tttaatgtga agcctgtttt atgaacagaa taaatgtggg 720  
 gtatattcag aataactaat gtttggaagt ttgttttatt ttgctaaaat tg 772

<210> 63  
 <211> 431  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 39  
 <223> 10-157-39 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 20..38  
 <223> 10-157-39.mis1

<220>  
 <221> misc\_binding  
 <222> 40..59

57

&lt;223&gt; 10-157-39.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 412..431

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 27..51

&lt;223&gt; 10-157-39 potential probe

&lt;400&gt; 63

ctgtgggkcc atttagtgat ttccctacct cccatcttyt attgcatcca caactgtggt	60
tctgtccata atttcctttg ctttctgtgc attattacat catatctgaa aatgagaaac	120
caaaaacaat agaaagcagc catgtctgga ggtgactggg gggtcgagaa gccctagttt	180
ctcaaaccct tagcaccaaa tttttccctc agttacactg agcgtttcac ttctgcagtg	240
atggagaagg gagatccctt atttcttctc atgagcatct ctggtgctgt ttcccttaga	300
gacaaataag gggttctatt taatgtgaag cctgttttat gaacagaata aatgtggtgt	360
atattcagaa taactaatgt ttggaagttt gttttatttt gctwaaaatt gggtctcaag	420
gcagctctgg t	431

&lt;210&gt; 64

&lt;211&gt; 431

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 131

&lt;223&gt; 10-157-131 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 111..130

&lt;223&gt; 10-157-131.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 132..150

&lt;223&gt; 10-157-131.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 412..431

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 119..143

&lt;223&gt; 10-157-131 potential probe

&lt;400&gt; 64

58

```

ctgtgggkcc atttagtgat ttccctacct cccatcttyt attgcatcca caactgtggt      60
tctgtccata atttcctttg ctttctgtgc attattacat catatctgaa aatgagaaac      120
caaaaacaat rgaaagcagc catgtctgga ggtgactggg gggtcgagaa gccctagtgt      180
ctcaaaccct tagcaccaaa tttttccctc agttacactg agcgtttcac ttctgcagtg      240
atggagaagg gagatccctt atttcttctc atgagcatct ctggtgctgt ttcccttaga      300
gacaaataag gggttctatt taatgtgaag cctgttttat gaacagaata aatgtggtgt      360
atattcagaa taactaatgt ttggaagttt gttttatatt gctwaaaatt ggttctcaag      420
gcagctctgg t                                                              431

```

&lt;210&gt; 65

&lt;211&gt; 431

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 166

&lt;223&gt; 10-157-166 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 146..165

&lt;223&gt; 10-157-166.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 167..186

&lt;223&gt; 10-157-166.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 412..431

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 154..178

&lt;223&gt; 10-157-166 potential probe

&lt;400&gt; 65

```

ctgtgggkcc atttagtgat ttccctacct cccatcttyt attgcatcca caactgtggt      60
tctgtccata atttcctttg ctttctgtgc attattacat catatctgaa aatgagaaac      120
caaaaacaat agaaagcagc catgtctgga ggtgactggg gggtcragaa gccctagtgt      180
ctcaaaccct tagcaccaaa tttttccctc agttacactg agcgtttcac ttctgcagtg      240
atggagaagg gagatccctt atttcttctc atgagcatct ctggtgctgt ttcccttaga      300
gacaaataag gggttctatt taatgtgaag cctgttttat gaacagaata aatgtggtgt      360
atattcagaa taactaatgt ttggaagttt gttttatatt gctwaaaatt ggttctcaag      420
gcagctctgg t                                                              431

```

&lt;210&gt; 66

&lt;211&gt; 431

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 246

&lt;223&gt; 10-157-246 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 226..245  
 <223> 10-157-246.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 247..266  
 <223> 10-157-246.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 412..431  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 234..258  
 <223> 10-157-246 potential probe

<400> 66							
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tctgtccata	atttcctttg	ctttctgtgc	attattacat	catatctgaa	aatgagaaac		120
caaaaacaat	agaaagcagc	catgtctgga	gggtgactgg	gggtcgagaa	gccctagttt		180
ctcaaaccct	tagcaccaaa	tttttccttc	agttacactg	agcgtttcac	ttctgcagtg		240
atggaraagg	gagatccctt	atctcttctc	atgagcatct	ctggtgctgt	ttcccttaga		300
gacaaataag	gggttctatt	taatgtgaag	cctgttttat	gaacagaata	aatgtgggtg		360
atattcagaa	taactaatgt	ttggaagttt	gttttatttt	gctwaaaatt	ggttctcaag		420
gcagctctgg	t						431

<210> 67  
 <211> 581  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 161  
 <223> 10-159-161 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 142..160  
 <223> 10-159-161.mis1

<220>  
 <221> misc\_binding  
 <222> 162..181  
 <223> 10-159-161.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..19  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind

<222> 403..422  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 149..173  
 <223> 10-159-161 potential probe

<400> 67  
 aggaattttt tttagggggg ttaatggtaa aggtgtttat atctgctaag gtaatttact 60  
 tgatatatgt ttgggttattt aagatatatg agttatgtta gctatttcat gtttaggctg 120  
 ctgtattttt agtaggctat attaaatatt tgaaaggatt wcattataaa gaacaaagtc 180  
 tcctaattctt tgatatagca ttgacatact ttttaaataat acaaggcata gaatatggcc 240  
 atttctgtta aatcatatat tcccaactgg ttattaatct aagaattcag aattttgagt 300  
 aattgctttt gcatcagatt atttacttca gtgctctcaa ttatgatggg gcattagaac 360  
 catctggggtt aacatttggt ttttattacc aatacctagg ctccaaccaa gtacagtga 420  
 actggaatgt acagagtgga caatggaacg aaggagaaca agaccaaagg acattttatt 480  
 tttatctgta tcagtgggtc aaagtccttt cagaaggagc atatagtgga cctaggtgat 540  
 tggtcrrmtt atccatcaaa gaggcacaca ccgaattagc a 581

<210> 68  
 <211> 581  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 162  
 <223> 10-159-162 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 142..161  
 <223> 10-159-162.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 163..182  
 <223> 10-159-162.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..19  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 403..422  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 150..174  
 <223> 10-159-162 potential probe

<400> 68  
 aggaattttt tttagggggg ttaatggtaa aggtgtttat atctgctaag gtaatttact 60  
 tgatatatgt ttgggttattt aagatatatg agttatgtta gctatttcat gtttaggctg 120  
 ctgtattttt agtaggctat attaaatatt tgaaaggatt tmattataaa gaacaaagtc 180  
 tcctaattctt tgatatagca ttgacatact ttttaaataat acaaggcata gaatatggcc 240  
 atttctgtta aatcatatat tcccaactgg ttattaatct aagaattcag aattttgagt 300  
 aattgctttt gcatcagatt atttacttca gtgctctcaa ttatgatggg gcattagaac 360  
 catctggggtt aacatttggt ttttattacc aatacctagg ctccaaccaa gtacagtga 420



61

actggaatgt	acagagtgga	caatggaacg	aaggagaaca	agaccaaagg	acattttatt	480
tttatctgta	tcagtgggtc	aaagtccttt	cagaaggagc	atatagtgga	cctaggtgat	540
tggtcrmttt	atccatcaaa	gaggcacaca	ccgaattagc	a		581

<210> 69  
 <211> 353  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 169  
 <223> 10-83-169 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 150..168  
 <223> 10-83-169.mis1

<220>  
 <221> misc\_binding  
 <222> 170..189  
 <223> 10-83-169.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..19  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 336..353  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 157..181  
 <223> 10-83-169 potential probe

<400> 69	
aagtcatact gcttcttact aggtctgtct ggggtgggaat gtaacttctt tggacctcaa	60
ttttcttatac tattgataaa agagattgga ctaggtgatt tccatcactt cttcccactc	120
tttgacttct ttataactta gtttgtctgt tttgctatct tcagggcayg accataataa	180
catccctgac ttctgtgctg cacaatgaca aagaattccc caaccagag atgtttgacc	240
ctggccactt tctggataag agtggcaact ttaagaaaag tgactacttc atgcctttct	300
cagcaggtaa tagatattca tttccatctg tccttcaggg cacatgatac ctt	353

<210> 70  
 <211> 425  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 152  
 <223> 10-84-152 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 132..151  
 <223> 10-84-152.mis1, potential

<220>

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<221> misc_binding
<222> 153..172
<223> 10-84-152.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 406..425
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 140..164
<223> 10-84-152 potential probe

<220>
<221> misc_feature
<222> 413
<223> n=a, g, c or t

<400> 70
gaacaaatcc cctatgtctc ttattttcag gaaaacggat gtgtatggga gagggcctgg      60
cccgcattgga gctgttttta ttcctgacca ccattttgca gaactttaac ctgaaatctc      120
aggttgaccc aaaggatatt gacatcacc cyattgcaa tgcatttggt cgtgtgccac      180
ccttgtagca gctctgcttc attcctgtct gaagaagggc agatagtttg gctgctcctg      240
tgctgtcacc tgcaattctc ccttatcagg gccattggcc tctcccttct ctctgtgagg      300
gatattttct ctgacttgtc aatccacatc ttcccatctc ctcaagatcc aatgaacatc      360
caacctccat taaagagagt ttcttgggtc acttcctaaa tatactctgct atnctccata      420
ctctg                                           425

<210> 71
<211> 425
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 243
<223> 10-84-243 : polymorphic base C or T

<220>
<221> misc_binding
<222> 224..242
<223> 10-84-243.mis1

<220>
<221> misc_binding
<222> 244..263
<223> 10-84-243.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 406..425
<223> downstream amplification primer, complement

```

<220>  
 <221> misc\_binding  
 <222> 231..255  
 <223> 10-84-243 potential probe

<220>  
 <221> misc\_feature  
 <222> 413  
 <223> n=a, g, c or t

<400> 71  
 gaacaaatcc cctatgtctc ttatttttcag gaaaacggat gtgtatggga gagggcctgg 60  
 cccgcacatgga gctgttttta ttccctgacca ccattttgca gaactttaac ctgaaatctc 120  
 aggttgaccc aaaggatatt gacatcaccc ccattgccaa tgcatttggc cgtgtgccac 180  
 ccttgatcca gctctgcttc attcctgtct gaagaaggcc agatagtttg gctgctcctg 240  
 tgytgtcacc tgcaattctc ccttatcagg gccattggcc tctcccttct ctctgtgagg 300  
 gatattttct ctgacttgtc aatccacatc ttcccattcc ctcaagatcc aatgaacatc 360  
 caacctccat taaagagagt ttcttgggtc acttcctaaa tatatctgct atnctccata 420  
 ctctg 425

<210> 72  
 <211> 425  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 277  
 <223> 10-84-277 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 257..276  
 <223> 10-84-277.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 278..297  
 <223> 10-84-277.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 406..425  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 265..289  
 <223> 10-84-277 potential probe

<220>  
 <221> misc\_feature  
 <222> 413  
 <223> n=a, g, c or t

<400> 72  
 gaacaaatcc cctatgtctc ttatttttcag gaaaacggat gtgtatggga gagggcctgg 60

64

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cccgcacgga gctgttttta ttctgacca ccattttgca gaactttaac ctgaaatctc 120
agggtgaccc aaaggatatt gacatcacc ccattgcaa tgcatttggc cgtgtgccac 180
ccttgtagca gctctgcttc attcctgtct gaagaagggc agatagtgtg gctgctcctg 240
tgctgtcacc tgcaattctc ccttatcagg gccattggc tctcccttct ctctgtgagg 300
gatattttct ctgacttgct aatccacatc tcccattcc ctcaagatcc aatgaacatc 360
caacctccat taaagagagt ttcttgggtc acttcctaaa tatatctgct atnctccata 420
ctctg 425

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<210> 73  
 <211> 425  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 295  
 <223> 10-84-295 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 275..294  
 <223> 10-84-295.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 296..314  
 <223> 10-84-295.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 406..425  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 283..307  
 <223> 10-84-295 potential probe

<220>  
 <221> misc\_feature  
 <222> 413  
 <223> n=a, g, c or t

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<400> 73
gaacaaatcc cctatgtctc ttattttcag gaaaacggat gtgtatggga gagggcctgg 60
cccgcacgga gctgttttta ttctgacca ccattttgca gaactttaac ctgaaatctc 120
agggtgaccc aaaggatatt gacatcacc ccattgcaa tgcatttggc cgtgtgccac 180
ccttgtagca gctctgcttc attcctgtct gaagaagggc agatagtgtg gctgctcctg 240
tgctgtcacc tgcaattctc ccttatcagg gccattggc tctcccttct ctctrtgagg 300
gatattttct ctgacttgct aatccacatc tcccattcc ctcaagatcc aatgaacatc 360
caacctccat taaagagagt ttcttgggtc acttcctaaa tatatctgct atnctccata 420
ctctg 425

```

<210> 74  
 <211> 424  
 <212> DNA  
 <213> Homo Sapiens

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<220>
<221> allele
<222> 43
<223> 10-85-43 : polymorphic base C or T

<220>
<221> misc_binding
<222> 24..42
<223> 10-85-43.mis1

<220>
<221> misc_binding
<222> 44..63
<223> 10-85-43.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..424
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 31..55
<223> 10-85-43 potential probe

<400> 74
gtttcttggg tcacttccta aatatatctg ctattctcca taytctgtat cacttgtatt      60
gaccaccaca tatgctaata cctatctact gctgagttgt cagtatgtta tcactagaaa      120
acaaagaaaa atgattaata aatgacaatt cagagccatt tattctctgc atgctctaga      180
taaaaatgat tattattttac tgggtcagtt cttagatttc tttcttttga gtaaaatgaa      240
agtaagaaat gaaagaaaat agaatgtgaa gaggtgtgac tggccctcat agtggttaagc      300
acaaaaaggg agaaaggtaa gagggtagga aagctgtttt agctaaatgc cacctagagt      360
tattggaggt ctgaatttgg aaaaaaaaaac tatgtccagg agcagctgta acvtgtaggg      420
aaat                                                                424

<210> 75
<211> 424
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 117
<223> 10-85-117 : polymorphic base G or T

<220>
<221> misc_binding
<222> 97..116
<223> 10-85-117.mis1, potential

<220>
<221> misc_binding
<222> 118..136
<223> 10-85-117.mis2, complement

<220>
<221> primer_bind
<222> 1..18

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<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..424
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 105..129
<223> 10-85-117 potential probe

<400> 75
gtttcttggg tcacttccta aatatatctg ctattctcca tactctgtat cacttgatt 60
gaccaccaca tatgctaata cctatctact gctgagttgt cagtatgta tcactakaaa 120
acaaagaaaa atgattaata aatgacaatt cagagccatt tattctctgc atgctctaga 180
taaaaatgat tattatttac tgggtcagtt cttagatttc tttcttttga gtaaaatgaa 240
agtaagaaat gaaagaaaat agaatgtgaa gaggctgtgc tggccctcat agtggttaagc 300
acaaaaaggg agaaaggtaa gagggtagga aagctgtttt agctaaatgc cacctagagt 360
tattggaggt ctgaatttgg aaaaaaaaaac tatgtccagg agcagctgta acvtgtaggg 420
aat 424

<210> 76
<211> 424
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 320
<223> 10-85-320 : polymorphic base A or T

<220>
<221> misc_binding
<222> 300..319
<223> 10-85-320.mis1, potential

<220>
<221> misc_binding
<222> 321..340
<223> 10-85-320.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..424
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 308..332
<223> 10-85-320 potential probe

<400> 76
gtttcttggg tcacttccta aatatatctg ctattctcca tactctgtat cacttgatt 60
gaccaccaca tatgctaata cctatctact gctgagttgt cagtatgta tcactagaaa 120
acaaagaaaa atgattaata aatgacaatt cagagccatt tattctctgc atgctctaga 180
taaaaatgat tattatttac tgggtcagtt cttagatttc tttcttttga gtaaaatgaa 240
agtaagaaat gaaagaaaat agaatgtgaa gaggctgtgc tggccctcat agtggttaagc 300

```

67

acaaaaaggg agaaaggtaw gagggtagga aagctgtttt agctaaatgc cacctagagt	360
tattggaggt ctgaatttgg aaaaaaaaaac tatgtccagg agcagctgta acvtgtaggg	420
aat	424

<210> 77  
 <211> 352  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 121  
 <223> 10-86-121 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 102..120  
 <223> 10-86-121.mis1

<220>  
 <221> misc\_binding  
 <222> 122..141  
 <223> 10-86-121.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 334..352  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 109..133  
 <223> 10-86-121 potential probe

<400> 77	
aagggagaaa ggthagaggg taggaaagct gttttagcta aatgccacct agagttattg	60
gaggtctgaa tttggaaaaa aaaactatgt ccaggagcag ctgtaacctg tagggaaata	120
mtggaacaat catccataag agggatgaac attaatgttt tgaattcatg ctctgctttt	180
gtgttactgt aaacacaaga tcaagatttg gataatcttt ttcctttgtg tttccaaactt	240
agatcatgtc taaatatatg ctttcatatg gctaatacatg tgtaaatgac tgttattttt	300
ctcttccaaa caagagcaaa atctccagaa atcctcacca ggctttattt tt	352

<210> 78  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-244-275 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-244-275.mis1, potential

<220>

<221> misc\_binding  
 <222> 502..521  
 <223> 12-244-275.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 228..247  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 660..678  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-244-275 potential probe

<400> 78  
 atcacgttct gtccagtgtc tgcctattcc cttcttcttt ttttcttccc ttgatgccct 60  
 tttatcacat gcattgtctc agacccttcg aatatgtgct cataaatgca tggcatcatc 120  
 tccttcccac atcgattcac tttcaattaa aagccaaaac tctttcattt caactttgga 180  
 tttaacatgc ttttgaaaga aggggtgaga aatatagaga aacagattgg gaaaccatgc 240  
 tctgctgttt ctttttttta aactttctat gtaagtgtgg aatttttcat tctgttttat 300  
 tattaacttt aagccaagac tttttaatag aaggatataat aaatacatct ttgtctatac 360  
 atttctgctg aatttgaaga aatgctgaat attcttaaac cattgtgttc cctggtgggc 420  
 tgatggactg tgattttata aggtggcctc agccaactgc agcagctgtt ccctgtcaga 480  
 ggggctagag gtttggcaag rgcggtggaa gaggtgcagt ggtgtgttcg ttcactagaa 540  
 gcatcagggg gaagggttttg cctgtttgta tttcatcttc tctcatcaag tcctcagaaa 600  
 ccacagtgtc gtctgcaggg tgctgtggat ctggcatggc ccatacaggc aacatgactg 660  
 agtagaaagg acacacagct ctggatgtcc ttgggccccca cagcaactgc ccttgaaca 720  
 tttagtccct gtgagcattt gatgatttac ttgccttcaa ttttccatgg acctaaact 780  
 ctttataaag ggaaatattt taaacctatg aaacattgtg gagaatggca tgggaaatac 840  
 ccattgtatgc accaccagc ttaacaaatg ctctcctgtc atttctaacc acaatctctt 900  
 tgaagagctc ttttgtcttt caatctctct tccctgtttg gccacatta cccttcatcg 960  
 tatgaagact tggatggctc ctgtgtcaga ctcttgtctg g 1001

<210> 79  
 <211> 950  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 450  
 <223> 12-251-153 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 431..449  
 <223> 12-251-153.mis1

<220>  
 <221> misc\_binding  
 <222> 451..470  
 <223> 12-251-153.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 298..318  
 <223> upstream amplification primer



<220>  
 <221> primer\_bind  
 <222> 806..826  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 438..462  
 <223> 12-251-153 potential probe

<220>  
 <221> misc\_feature  
 <222> 52,54,100  
 <223> n=a, g, c or t

<400> 79  
 taggaagctg agagaaaagg atcttctccc ctgtgcacag gagccaggaa tnanttatct 60  
 gttacattgt ccccttggat attccaaaag aataatacan tgtgtagaaa gaaaaataaa 120  
 aaacctgatt gtactaaagt tttgagcata gacgttatgg agtggtagga ggggaagggtg 180  
 tttgatagct gttggctggc agtgactggg gcaggaaagt tacaatgagg aagttggaat 240  
 aacttcaatc ccttcattat tttgtgagg ataccaccaa aatatgaaat attaaacctt 300  
 cccaccactt ctaatttctt ttctccaatc ttaaatttta aaagactctg taaaggctat 360  
 aggtagggca atgctattgt ttgttgtctg aactggaatg taatttgaat tatgctggaa 420  
 caacatgtgt aagctgagcc tgtcctgggm agactgggga catgtggtca ctcagctatg 480  
 ggatgccccg atcaactttg gagtgatcta tttgtttaat caatatcact gttagtctt 540  
 acttttacaa aaataatctg ccctcagagc aacctcagat cccaggagtt ggggaaaagc 600  
 aggtgtttca ggaggcttat caggagtgagc agcggagaca tgacgttcac agcaagtctg 660  
 aacaggggtg ggctgttcta taaagtgtg aagagacatc agctctgggc gtagactgtg 720  
 gggctctggca atgtcaaatt tattgattgg ccgagaaaga gttaattatt ttattcttgt 780  
 cctgcagaag cacagtgttg acacaccttt taccatccac actcaacaca aactactgta 840  
 attgtctgat tattgggtct gtgtctccct atgactgagg tccttgagct cagaggtggg 900  
 tctaactcac cttagtgtct ccatcactcc cagcacaggg ccagctgcat 950

<210> 80  
 <211> 746  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 246  
 <223> 12-254-115 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 226..245  
 <223> 12-254-115.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 247..266  
 <223> 12-254-115.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 132..152  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 586..603  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 234..258  
 <223> 12-254-115 potential probe

```
<400> 80
ctgccacaat ctcttgagtc cagacaatct agagcagaag ggtagactga ggaaaatata      60
cacagtataa aaaagtaaca aaatcaaaac ctgaaacaaa gatcaacatc caataaatgc      120
ttctgaataa agggagagta gataagaact ggattttaat cccaacactg ccattttacca      180
gctggccaat actgagctag ttactctaaa gagttcagtt ttctcatttg tacaaatagg      240
atttgwcttt ccattctcact gagttgtgat gagagtcata tgcaacagca tatgaagagg      300
ctagcaaaaag gtattttaaca agcgttcaac attctcatga tgacatgaat aacactgtac      360
atacaacata ccaacttgat aaatacacag cacagttaat agctgagggc agagtttatgg      420
ttgggaagag agagagtgca acataggcag agtgaggggg gattcccaca atttttctaag      480
acagaaaagt gggggaatca gtagttactg gaaagaatag gcaatgcctg actggataga      540
aaaagattct atgcctttgt caaatttcac aaaagtgact taagcctata ctgcgggatg      600
ttcacactac gtcccttttag tgcagttacg gtacttcagg ctgcaagtaa ccaaatacaa      660
ctaaaattgt cttataacaat aagggcgtaa ttatctcata taacaagaag cttggcatga      720
aggaaatttc aacaatttca caacgg                                     746
```

<210> 81  
 <211> 811  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 311  
 <223> 12-254-180 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 292..310  
 <223> 12-254-180.mis1

<220>  
 <221> misc\_binding  
 <222> 312..331  
 <223> 12-254-180.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 132..152  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 586..603  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 299..323  
 <223> 12-254-180 potential probe

```
<400> 81
ctgccacaat ctcttgagtc cagacaatct agagcagaag ggtagactga ggaaaatata      60
cacagtataa aaaagtaaca aaatcaaaac ctgaaacaaa gatcaacatc caataaatgc      120
ttctgaataa agggagagta gataagaact ggattttaat cccaacactg ccattttacca      180
gctggccaat actgagctag ttactctaaa gagttcagtt ttctcatttg tacaaatagg      240
atttgctttt ccattctcact gagttgtgat gagagtcata tgcaacagca tatgaagagg      300
ctagcaaaaag rtattttaaca agcgttcaac attctcatga tgacatgaat aacactgtac      360
atacaacata ccaacttgat aaatacacag cacagttaat agctgagggc agagtttatgg      420
```

71

ttgggaagag agagagtgca acatagggcag agtgaggggg gattcccaca attttctaag	480
acagaaaaagt gggggaatca gtagttactg gaaagaatag gcaatgcctg actggataga	540
aaaagattct atgcctttgt caaatttcac aaaagtgact taagcctata ctgcgggatg	600
ttcacactac gtcccttttag tgcagttacg gtacttcagg ctgcaagtaa ccaaatacaa	660
ctaaaaattgt cttatacaat aagggcgtaa ttatctcata taacaagaag cttggcatga	720
aggaaaatttc aacaatttca caacggcaac aaaaactctg tttcttctac ctttccacca	780
ttcctgtgtt ctcagttcca atatggctgc t	811

&lt;210&gt; 82

&lt;211&gt; 999

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-265-300 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-265-300.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-265-300.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 779..798

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 308..328

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-265-300 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 427,494,669..670,679,772,874,944

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 82

taataatgtg ttttggggta agcctactca tattctcaac ctgtctgcag tagtcgttag	60
aatctgaact tcctgaagtt catgtgcaaa gttgagttaa ttgtttaata ttcaacaagg	120
attatgccag taagatggta ggaaaatatt agatatgtgt catcactgct ggtattattt	180
aaactgcaac atattttagc tggctgctga tctcagccac catgcctgca ttttatctct	240
gtctcgtggt ctgcaacctt ggaagctttg aacttagctc atagaatcct gggcatcaag	300
aacatgtggt tctaattggct agatagggaa tgagagtaaa aggattttgc ccacggtcac	360
gtgagtaaac aacagatttg gaggggtctg gactactgtg atgacttcat tctgacaata	420
tgttccnagt tgtcctttca tttcctccta atcacatgtc tggctctgatc tggctgtttc	480
ccaccttcca attncctgyc ttctccaatg ctcccttccg taggtcactc tgtggctcag	540
agaccctgct tagcaagcgc ccaacctttc aattattttgt tcagtaaaac ttgaactcat	600
gtctcccttt cttgataaaa agaaaatacg ttatgtaatg tcgggttact ctataactct	660
tgtctcgtgn ctctcggnna actactgaac taactgtttt catattgagc aaacgtttat	720
ggaaggactg ccaagagtca ggtactaggc ttggtaatat tccccgttct cntctagtca	780
aagccaacac cagccagact tgcagatcta ggtcccaagc ccactgcaga tcacaggcca	840

72

```

gggtctggtc tcctctgagc tcctttggga gggnaaagac agaattatta acacccattt 900
tgtagattag gcaactgagg ctgaggaagt ttaaataact cagnacaggg cctgcacgtc 960
agtcattatc caaggatccc tactcactgt cttctctct 999

```

```

<210> 83
<211> 1001
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele
<222> 501
<223> 12-271-118 : polymorphic base T or C

```

```

<220>
<221> misc_binding
<222> 481..500
<223> 12-271-118.mis1, potential

```

```

<220>
<221> misc_binding
<222> 502..521
<223> 12-271-118.mis2, potential complement

```

```

<220>
<221> primer_bind
<222> 598..618
<223> upstream amplification primer, complement

```

```

<220>
<221> primer_bind
<222> 122..141
<223> downstream amplification primer

```

```

<220>
<221> misc_binding
<222> 489..513
<223> 12-271-118 potential probe

```

```

<220>
<221> misc_feature
<222> 205,525,542,571,657
<223> n=a, g, c or t

```

```

<400> 83
gactgagtcc gaaaaggagt ctgcaaaggg agataggggt gggtcagttt tataggactg 60
gggtaagcag tggaaagttg cagttaaagg aagttatcta ttgtcagcag aggagggggg 120
cacaagggtg atggtagggg gatcataaga ctcattgtcc agaagaagaa tgtcacgagg 180
tcgatcaatc gatcagttgg ggcangggca gtaacaagtc ataatggaac gttgtaaggt 240
tggccaatca gttaagacag gagctggctg tttcacctat ttgtagtttt tggttgcctc 300
aggccatctg gatgtaccca tgcaggcttg ggctaagagg cctgaaaccc accactttcc 360
catgtcaaat ctttagtaga tgtaccccca agatacacat tcctcggacc ttcttttcca 420
tagttaaaac ttcacccctg aaatgtagaa acaggaaggt ttttttttta agtttcagt 480
caaactcgga gcaagtgtca yaattttctg tctccgatgt gtagnagggt acattttctc 540
angaactttc acgttaagct ggaaaactgg naaagcgagt ccactttgtc attctgtcac 600
tcaactcatt tctcactcaa caaacatgcc tcacacttat ctaaatctgc tagactnaaa 660
agagggtccct ggtgtctgta acttttcta tctgctagaa ttctagagt agctcatgaa 720
ataaatgaaa aggatgaaga acaaagagaa aaaagactgc acgttccctt ctggcgctca 780
ctcacattcc ctcagcctca gttttctccac atgcccctag aggtgatcat tcaaggattt 840
atgagatttt agagacaaca catgaaaaag caaagagaca tcagaaagac aaggagttac 900
ttagtattta tacacaagga taagacattc agtatcgaca acacttaaag aaaattcaag 960
agtgtattta aatttcccat ttcaaatacc tcctctattt t 1001

```

73

<210> 84  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-272-112 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-272-112.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-272-112.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 390..409  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 768..788  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-272-112 potential probe

<400> 84  
 aagagggggcc caacttgtaa tcataggagc ttatgctatt ttaatgccat ccatcagact 60  
 acaatcaatt accactcatc tagctttttg tccatctctc attcttgtag atcctgagat 120  
 agtcaattct gagaactgta gcctagatct atcacctgat gcctctcaaa gatataatcc 180  
 gtgcttctca agctaggcta tgcacacaaa tcaactgcac ttgtgaaagt tcagattttg 240  
 aatcagtagt tcaaggggtg ggtttgagat ttgcatctc taatgagctc tcagatgctt 300  
 ctgacccatg gaccacactt tgaataccaa gaagtggctc gtagaccaat attggtccct 360  
 taagtccctc caaacatata ttcgggaaac gtcctttgat tttccctaca tttaaccatt 420  
 agtggtgcaa attctctcaa agtttgctca gatataattg agctaaaata aattacattt 480  
 ttcttggggg agagtactac mtcataattaa cttacaataa agtactttta ggatcattca 540  
 aggaacacac ccataacact gagtatgtta tgcggaaatg ctctctctgg aaattacaca 600  
 gctgtgcagg tggcgggggt ggcatgagga ggagtggatg gccacattc tcgaagacct 660  
 tgggggaaaac tggattaaaa tgatttgctt tattctgggt ctgtaagata cacatcagaa 720  
 tgaaaccacc ccagtgtag ctctgaattg cttttctatt cttttccctt agggatttga 780  
 gggcttcaact tagatttctc ttcattctaaa ctgtgatgcc ctacattgat ctgatttacc 840  
 taaaatgtct ttcctctcct ttcagctctg tccgatctgg agctcgtggc ccaatcaatt 900  
 atctttatct ttgctggcta tgaaaccacg agcagtggtc tctccttcat tatgtatgaa 960  
 ctggccactc accctgatgt ccagcagaaa ctgcaggagg a 1001

<210> 85  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 10-216-182 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 10-216-182.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..522  
 <223> 10-216-182.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 323..339  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 800..819  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 10-216-182 potential probe

<220>  
 <221> misc\_feature  
 <222> 76,80,96,319,399,556,578,667,676,731,752,759,914,933  
 <223> n=a, g, c or t

<400> 85  
 tgctatcctt cccccctctc cccacccac aacaggcccc agtgtgtgat gttccccttc 60  
 ctgtgtccat gtgttnctcn attgtatatt tttttnaaat ctaccacatc aaggcacctc 120  
 tttttcatgt tgcccattgt ttaggtgaac ataaagacag agctcgtctg aggcaacata 180  
 cagtccaaca aagccacctg cctctctgtc tccactctct ctctacactg cacgcgtgct 240  
 aggtgttgat cctgtctatt ccagtgggaag aacagggtcc gtaccatgtg gagaatttgc 300  
 atgtaaaagg agactgggna tatacaggct ggagaccaca tcaggtggct gggcatgtgg 360  
 gataaatcct attgagcatc tgtcataggg cctgtcacnt tagtagacag tcactaaata 420  
 tttgttaaat acatgatgcc tgtttaacac attttctaca accatggaga cctccacaac 480  
 tgatgttaga caaaatcttt ctrctttgaa ctctagcctt tcgggccagt gggatttatg 540  
 aaaaatgcc a tctctnatag ctgaggatga agaattggnaa gagaatacga tcattgctgt 600  
 ctccaacatt caccagcgga aaactcaagg aggtatgaaa ataacttggg ttttaattag 660  
 aaacttnaaa gaatgnaatc aggtggggac aggtagaaa taagatcaga gttcctttcc 720  
 gaggagtagt nctgctgaat ttgagcttcc tnaaaaatna gtctttttat gtacagaaaa 780  
 cacatcataa aattcattac acaatgtcac ttattgttcc atgccaggca aagtcattgtc 840  
 cttctgggac ttatgtctgc acatttaact atgggtgggtg ttgtgttttg tgcttagatg 900  
 gtccctatca ttgncccagt atggagatgt gtntgggtgag aaatctgagg cgggaagcag 960  
 agacaggcaa gcctgtcacc ttgaaacagt aagtaggagc a 1001

<210> 86  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-217-91 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 482..500

<223> 10-217-91.mis1

<220>

<221> misc\_binding

<222> 502..521

<223> 10-217-91.mis2, potential complement

<220>

<221> primer\_bind

<222> 411..427

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 761..777

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 10-217-91 potential probe

<220>

<221> misc\_feature

<222> 20,100,257,279,368,377,432,453,460,615,634,921,946,986

<223> n=a, g, c or t

<400> 86

catgtaaaag gagactgggn atatacaggc tggagaccac atcaggtggc tgggcatgtg	60
ggataaatcc tattgagcat ctgtcatagg gcctgtcacn ttagtagaca gtcactaaat	120
atttgttaaa tacatgatgc ctgtttaaca cattttctac aaccatggag acctccacaa	180
ctgatgtagg acaaaatctt tctgctttga actctagcct ttcggggccag tgggatttat	240
gaaaaatgcc atctctnata gctgaggatg aagaatggna agagaatacg atcattgctg	300
tctccaacat tcaccagcgg aaaactcaag gaggtatgaa aataacttgg gttttaatta	360
gaaacttnaa agaatgnaat caggtgggga caggtagaaa gtaagatcag agttcctttc	420
cgaggagtag tnctgctgaa tttgagcttc ctnaaaaatn agtcttttta tgtacagaaa	480
acacatcata aaattcatta yacaatgtca cttattgttc catgccaggc aaagtcatgt	540
ccttctggga cttatgtctg cacatttaac tatgggtggg gttgtgtttt gtgcttagat	600
ggtccttacc attgncccag tatggagatg tgtntggatg gaaatctgag gcggaagca	660
gagacaggca agcctgtcac cttgaaacag taagtaggag cacagccatg gggttctgag	720
ctgtcatgag cccctccagc tgcctgctat ggagctgata ctcccgtgt tgggttattc	780
cagtgaccag acaaaaggag ggctgtggta atgcaacttc aatgggtctc ccaagatggg	840
gcagctccga tgaggagggtg gggcagctgg aggaaaagga tcttctcccc tgtgcacaga	900
ggtcagggtt tacatatctg nttaaattgt caccttgatg attctnggag gactaaatac	960
atcctttagg gggaaaagtg tgattngtat caaagtttta a	1001

<210> 87

<211> 1001

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 503

<223> 10-213-292 : polymorphic base G or C

<220>

<221> misc\_binding

<222> 484..502

<223> 10-213-292.mis1

<220>

<221> misc\_binding

<222> 504..523  
 <223> 10-213-292.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 212..230  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 590..608  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 10-213-292 potential probe

<220>  
 <221> misc\_feature  
 <222> 86,281,718,743,788,878,885,890,1000  
 <223> n=a, g, c or t

<400> 87  
 attagtcattg tttgcttcca tgagaaagaa aaaccactac atgggttatgc taaggatttc 60  
 agtcattggg gtttagagcct tcccgnaatg tctcctgctt tcataactcc tccacacatc 120  
 ttagtggggc attgagcaca tcaaagggca tgacagttat taaaatactt tatgaatgct 180  
 acaatccttt gccagtatga gttgttctct ggaacttcta acagttcaac agtactacat 240  
 ggactgagtt aaaagttaat tcaaaaatct caatttatcc naaatctggt tctttctttt 300  
 caggcaccac ccacctatga tactgtgcta cagttggagt atcttgacat ggtggggaat 360  
 gaaacactca gattattccc agttgctatg agacttgaga gggctctgcaa aaaagatggt 420  
 gaaatcaatg ggatgtttat tcccaaaggg gtgggtggga tgattccaag ctatgttctt 480  
 catcatgacc caaagtactg gasagagcct gagaagttcc tccctgaaag gtaggaggcc 540  
 cctgggaagg gagccctccc tgaaccagcc tgggtcaagc atattctgcc tctctacagg 600  
 acagtctggg cttgtacaat catttgcttg tctttttatg tttaaaagg tttttcaaat 660  
 catgaaattg atcattgtca cactttacaa accacagact agataaaaaga aaactatnag 720  
 ccagtcacag tcccagcaac ttnaagatga aggtcctcaa ttatgtcctt atgggtcata 780  
 agtgtccnaa aatgtaagga ctctttttaa aacacatgat cacaatgcta ttattatgtc 840  
 ccacaaatga atattttttc ctgaatataa tcaaatacnt caggnaatcn aaatttgaat 900  
 aaaaaacatg cgtctaattc tcaaagaatt tatagggttag tgcaacagat agacaaagaa 960  
 agcagtgatg acactgcttt ccatcaatac agtagcatcn a 1001

<210> 88  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-214-279 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-214-279.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-214-279.mis2, potential complement

<220>



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<221> primer_bind
<222> 154..174
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 746..763
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-214-279 potential probe

<400> 88
ctcatccatt ttacttaaaa tttaaatacaaaa aaaagaacac aggtttccat gaatttgctct 60
caggcctggc acagaatagt actccataaa tattttgtta aatgatatag gatgaatgct 120
ctcactgtcc aatcttcaca catcttatag actaagtata aagaatccaa gattttatagt 180
gctgaaagta gtttttatat gtttacaaag cattattgtc attactgcat tttttttgcc 240
cattactcca tagagatcag aatatcactc tgttggtgcc cctcaacact gaaggagtgt 300
ctcactcact ttgatgctat actttctact tttgtttatt taatgcttct caatatgctt 360
gtttaactgt tgcagatccc cctgaaaatta agcttaggag gacttcttca accagaaaaa 420
cccgttggtc taaagggtga gtcaagggat ggcaccgtaa gtggagcctg aattttccta 480
aggacttctg ctttgctctt yaagaaatct gtgcctgaga acaccagaga cctcaaatta 540
ctttgtgaat agaactctga aatgaagatg ggcttcaccc aatggactgc ataaataacc 600
ggggattctg tacatgcatt gagctctctc attgtctgtg tagagtgtta tacttgggaa 660
tataaaggag gtgaccaaat cagtgtgagg aggtagattt ggctcctctg cttctcacgg 720
gactatttcc accaccccca gttagcacca ttaactcctc ctgagctctg ataagagaat 780
caacatttct caataatttc ctccacaaat tattaatgaa aataagaatt attttgatgg 840
ctctaacaat gacatttata tcacatgttt tctctggagt attctataag ttttatgtta 900
aatcaataaa gaccacttta caaaagtatt atcagatgct ttcctgcaca ttaaggagaa 960
atctatagaa ctgaatgaga accaacaagt aaatatTTTT g 1001

<210> 89
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-214-380 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 10-214-380.mis1, potential

<220>
<221> misc_binding
<222> 502..520
<223> 10-214-380.mis2, complement

<220>
<221> primer_bind
<222> 124..143
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 647..664
<223> downstream amplification primer, complement

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<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-214-380 potential probe

<220>  
 <221> misc\_feature  
 <222> 398,723,730,789  
 <223> n=a, g, c or t

<400> 89  
 gaacacgggt tcccatgaat ttgtctcagg tcaaacctca cacagaatag gtgctccatg 60  
 aatattttgt taaatgatag atgatgaatg ttctcactat ccaatcttca cacatcttat 120  
 agagtaagta taacgaatcc aagatttata gtgctgaaag tagtttttat atgtttacaa 180  
 agcattattg tcagtaatgt ttttttactt tgatgctata ctttctactt ttgctttatt 240  
 taatgcttct caatatgctc gtttaactgt tgcagatccc cctgaaatta cgctttggag 300  
 gacttcttct aacagaaaaa cccattgttc taaaggctga gtcaagggat gagaccgtaa 360  
 gtggagcctg atttccctaa ggacttctgg tttgctcntt taagaaagct gtgccccaga 420  
 acaccagaga cctcaaatta ctttacaat agaacctga aatgaagacg ggcttcaccc 480  
 aatgtgctgc ataaataatc rgggattctg tacgtgcatt gtgctctctc atggtctgta 540  
 tagagtgtta tacttggtta tatagaggag atgaccaa atcagtgctggg gaagtagatt 600  
 tggcttctct gcttctcata ggactatctc caccacccc agttagcacc attaaactct 660  
 cctgagctct gataacataa ttaacatttc tcaataattt caaccacaat cattaataaa 720  
 aantaggaan ttattttgat ggctctaaca gtgacattta tatcatgtgt tatatctgta 780  
 gtattctant agtaagcttt atattaagca aatcaataaa aacctcttta caaaagtatt 840  
 attggatgtt tcctgcacat taaggagaaa tctatagaac tgaatgactg agaaccaaca 900  
 actaaatatt ttgatcattg taatcactgt tgggtgtggga actggagtgc agtggtgcaa 960  
 tcttggctca ctgcgagctc tgcctcccag gttcacgcca t 1001

<210> 90  
 <211> 437  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 216  
 <223> 2-1-216 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 196..215  
 <223> 2-1-216.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 217..236  
 <223> 2-1-216.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 417..437  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 204..228  
 <223> 2-1-216 potential probe

<400> 90  
gccccaaagaa gtgtgttgtg tttgcttatt tcttacagag taatgctgaa atctgtgttg 60  
cttttcccca ccagggtcatt atcagtagca gaagtgttc ctgggtcatg agtcgggtct 120  
gggatgatgg ctatccttgg gatatgatgt atgttaccg ctttgcaccc tttctccgga 180  
atgtccttcc ttcattcatc tctgactggt tataatrtcca gaagatgaac acgtgggtta 240  
agcatgagaa ctatggcctg atgcctttaa atgggtactt aaaaatggaa atttttttta 300  
ttcaaaaaag gggggcactc atttaataa tttattctct ctagaactta cttttgttgt 360  
ctcattgagc ctagaacat taaactcaag gtttcatagg tgacggaata tgcccagaga 420  
ccacgtatgg cttggaa 437

<210> 91  
<211> 437  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 397  
<223> 2-1-397 : polymorphic base C or T

<220>  
<221> misc\_binding  
<222> 377..396  
<223> 2-1-397.mis1, potential

<220>  
<221> misc\_binding  
<222> 398..417  
<223> 2-1-397.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 1..18  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 417..437  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 385..409  
<223> 2-1-397 potential probe

<400> 91  
gccccaaagaa gtgtgttgtg tttgcttatt tcttacagag taatgctgaa atctgtgttg 60  
cttttcccca ccagggtcatt atcagtagca gaagtgttc ctgggtcatg agtcgggtct 120  
gggatgatgg ctatccttgg gatatgatgt atgttaccg ctttgcaccc tttctccgga 180  
atgtccttcc ttcattcatc tctgactggt tataatrtcca gaagatgaac acgtgggtta 240  
agcatgagaa ctatggcctg atgcctttaa atgggtactt aaaaatggaa atttttttta 300  
ttcaaaaaag gggggcactc atttaataa tttattctct ctagaactta cttttgttgt 360  
ctcattgagc ctagaacat taaactcaag gtttcayagg tgacggaata tgcccagaga 420  
ccacgtatgg cttggaa 437

<210> 92  
<211> 426  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele

80

<222> 232  
 <223> 2-3-232 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 212..231  
 <223> 2-3-232.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 233..252  
 <223> 2-3-232.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 406..426  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 220..244  
 <223> 2-3-232 potential probe

<220>  
 <221> misc\_feature  
 <222> 419  
 <223> n=a, g, c or t

<400> 92  
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 agccagccgt gttcacagtg tcaaatgaag ggatgtcttt gattgcttcc aggtgttcc 120  
 cagcaccacc ggagggggat gggatgatcag ccgaatcttt gactcgggct acccatggga 180  
 catgggtgttc atgacacgct ttcagaacat gttgagaaat tccctcccaa cyccaattgt 240  
 gacttggttg atggagcgaa agataaaciaa ctggctcaat catgcaaatt acggcttaat 300  
 accagaagac aggtaaaatat aatgtgactg ccaagggctt ttaggaagaa ggagcctctg 360  
 cctgtccagc agcctataca agccaggcag taccacagca acatggctga atgtgtggna 420  
 acactt 426

<210> 93  
 <211> 429  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 51  
 <223> 2-4-51 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 31..50  
 <223> 2-4-51.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 52..71  
 <223> 2-4-51.mis2, potential complement

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<220>
<221> primer_bind
<222> 3..24
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..429
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 39..63
<223> 2-4-51 potential probe

<400> 93
gygctgttac tgtaaagaca ttgcattact actgttgacc tcagagcacg mgcctcttgc      60
ctaattctag gactcctaac taagtctttg gagtttcagc tggaagaatg ctggaggaat      120
acggaactcc tcccatctct cacagccacc tccaactctt aaaaacgctt ccaactgcct      180
cccagcacac aaccaaggga gaaaactatt ctgtcaaaga gacggtgcca aaaggcaaaa      240
acaaaggtaa ggatgatcgc tggggaaaga agctgaaaag gaaaagctca gaactctagc      300
tggaattttg gctcacatcc ctagtatgtt actgcatagt ctggctttgt tcaatgggtc      360
gcttttaaat attaaagcta gatgtaagca aggtttgcaa caaagtccat aagaaactca      420
gcttttctc                                     429

<210> 94
<211> 429
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 126
<223> 2-4-126 : polymorphic base A or G

<220>
<221> misc_binding
<222> 106..125
<223> 2-4-126.mis1, potential

<220>
<221> misc_binding
<222> 127..146
<223> 2-4-126.mis2, potential complement

<220>
<221> primer_bind
<222> 3..24
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..429
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 114..138
<223> 2-4-126 potential probe

<400> 94
gygctgttac tgtaaagaca ttgcattact actgttgacc tcagagcacg agcctcttgc      60
ctaattctag gactcctaac taagtctttg gagtttcagc tggaagaatg ctggaggaat      120

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82

acggarctcc	tcccatttct	cacagccacc	tccaactctt	aaaaacgctt	ccaactgcct	180
cccagcacac	aaccaagggg	gaaaactatt	ctgtcaaaga	gacggtgcc	aaaggcaaaa	240
acaaaggtaa	ggatgatcgc	tggggaaaga	agctgaaaag	gaaaagctca	gaactctagc	300
tggaaatttg	gctcacatcc	ctagtatggt	actgcatagt	ctggctttgt	tcaatgggtc	360
gcttttaaat	attaaagcta	gatgtaagca	aggtttgcaa	caaagtccat	aagaaactca	420
gcttttctc						429

&lt;210&gt; 95

&lt;211&gt; 420

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 202

&lt;223&gt; 2-5-202 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 182..201

&lt;223&gt; 2-5-202.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 203..222

&lt;223&gt; 2-5-202.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..25

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 400..420

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 190..214

&lt;223&gt; 2-5-202 potential probe

&lt;400&gt; 95

tgtttaaagc	caatthcctg	agcacatcat	aaggattctc	ttaccggttg	tcccagttaa	60
gtaatgttga	ttgatcaact	ccttgacagg	agctgatggc	aaagaaggta	gctgtgattg	120
gagctggggt	cagtggccta	atttctctga	agtgtgtgtg	ggatgaggga	cttgagccca	180
cttgctttga	gagaactgaa	grtattggag	gagtgtggag	gttcaaagta	agtgagattt	240
tcttgggtct	tgaacagggt	gtgttggtat	ttcagggtga	atcacagtta	ctgatgggtc	300
atattgagaa	atttattaaa	caactctgat	cagattttat	ttctacttat	tgatgtggcc	360
ataatggaac	tgaagtcata	ggctggcatc	tctccccag	tcaataactaa	cccaaccag	420

&lt;210&gt; 96

&lt;211&gt; 420

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 275

&lt;223&gt; 2-5-275 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

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<222> 255..274
<223> 2-5-275.mis1, potential

<220>
<221> misc_binding
<222> 276..295
<223> 2-5-275.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 400..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 263..287
<223> 2-5-275 potential probe

<400> 96
tgtttaaaagc caatthcctg agcacatcat aaggattctc ttaccggttg tcccaggttaa      60
gtaatgttga ttgatcaact ccttgacagg agctgatggc aaagaaggta gctgtgattg      120
gagctggggt cagtggccta atttctctga agtgcgtgtg ggatgaggga cttgagccca      180
cttgctttga gagaactgaa gatattggag gagtgtggag gttcaaagta agtgagattt      240
tcttgggtct tgaacagggt gtgttggtat ttcarggtga atcacagtta ctgatgggtc      300
atattgagaa atttattaaa caactctgat cagattttat ttctacttat tgatgtggcc      360
ataatggaac tgaagtcata ggctggcatc tctcccccag tcaataactaa cccaaccagg      420

<210> 97
<211> 420
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 346
<223> 2-5-346 : polymorphic base C or T

<220>
<221> misc_binding
<222> 326..345
<223> 2-5-346.mis1, potential

<220>
<221> misc_binding
<222> 347..366
<223> 2-5-346.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 400..420
<223> downstream amplification primer, complement

<220>

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<221> misc\_binding

<222> 334..358

<223> 2-5-346 potential probe

<400> 97

tgttttaaagc caatthcctg agcacatcat aaggattctc ttaccgggtg tcccagttaa	60
gtaatgttga ttgatcaact ccttgacagg agctgatggc aaagaaggta gctgtgattg	120
gagctgggggt cagtggccta atttctctga agtgctgtgt ggatgaggga cttgagccca	180
cttgctttga gagaactgaa gatattggag gagtgtggag gttcaaagta agtgagattt	240
tcttgggtct tgaacagggt gtgttggtat ttcagggtga atcacagtta ctgatgggtc	300
atattgagaa atttattaaa caactctgat cagattttat ttctayttat tgatgtggcc	360
ataatggaac tgaagtcata ggctggcatc tctccccag tcaataactaa cccaaccag	420

<210> 98

<211> 430

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 171

<223> 2-8-171 : polymorphic base A or G

<220>

<221> misc\_binding

<222> 151..170

<223> 2-8-171.mis1, potential

<220>

<221> misc\_binding

<222> 172..191

<223> 2-8-171.mis2, potential complement

<220>

<221> primer\_bind

<222> 1..18

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 405..427

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 159..183

<223> 2-8-171 potential probe

<400> 98

ggttcaagat ycctcagcaa atgaccttc agaatgtttt tcttctgtat gtctcagata	60
cattatgaag gaacctgtac taaatgatga tgtcccaagt cgtctactct gtggagccat	120
caaggtgaaa tctacagtga aagagctcac agaaacttct gccatctttg rggatggaac	180
agtggaggag aacattgatg tcatcatttt tgcaacagga tatagtttct cttttccctt	240
ccttgaagat tcaactcgta aagtagagaa taatatggtc tcaactgtata aatacatatt	300
ccccgctcac ctggacaagt caaccctcgc gtgcattggc ctcacccagc ccctagggtc	360
cattttccca actgctgaac ttcaagctcg ttgggtgaca agagtttcaa aggtaagtgt	420
gtaggcaggt	430

<210> 99

<211> 428

<212> DNA

<213> Homo Sapiens



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<220>
<221> allele
<222> 188
<223> 2-9-188 : polymorphic base A or G

<220>
<221> misc_binding
<222> 168..187
<223> 2-9-188.mis1, potential

<220>
<221> misc_binding
<222> 189..208
<223> 2-9-188.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 176..200
<223> 2-9-188 potential probe

<400> 99
tcacatwgag tgctatgggg gtggcacccc ctgaagtcca acagcacgga agccctgact      60
ggtatgacat ggttcaatgt ccagagtcca attttaagaa tcaacaacta gacaaagtaa      120
tgatattgac tcaaaacttac tattcaaacc aaccttttat tccttaggct tgtgtagcct      180
gccctcarag agaactatga tgatggacat tatcaaaagg aatgaaaaaa gaattgacct      240
gtaagaattt tttttaattc tttacatgaa gcagtgtttc tcaaagtaca gtgatctaac      300
tacttacaag aaccacctag ctgcctgata aaatgcaaat ttctgggcta tagcccagat      360
gattgaatca gaaactccgt gtgtgaggct aaaaagttgc atttttatct tcttcctaag      420
ggcatag                                           428

<210> 100
<211> 428
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 223
<223> 2-9-223 : polymorphic base G or T

<220>
<221> misc_binding
<222> 203..222
<223> 2-9-223.mis1, potential

<220>
<221> misc_binding
<222> 224..243
<223> 2-9-223.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21

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<223> upstream amplification primer

<220>
<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 211..235
<223> 2-9-223 potential probe

<400> 100
tcacatwgag tgctatgggg gtggcacccc ctgaagttca acagcacgga agccctgact      60
ggatgacat gggtcaatgt ccagagttta attttaagaa tcaacaacta gacaaagtaa      120
tgatattgac tcaaacttac tattcaaacc aaccttttat tccttaggct tgtgtagcct      180
gccctcagag agaactatga tgatggacat tatcaaaagg aakgaaaaaa gaattgacct      240
gtaagaattt tttttaattc ttacatgaa gcagtgttc tcaaagtaca gtgatctaac      300
tacttacaag aaccacctag ctgcctgata aaatgcaaat ttctgggcta tagcccagat      360
gattgaatca gaaactccgt gtgtgaggct aaaaagttgc atttttatct tcttcctaag      420
ggcatag                                           428

<210> 101
<211> 450
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 107
<223> 2-10-107 : polymorphic base C or T

<220>
<221> misc_binding
<222> 87..106
<223> 2-10-107.mis1, potential

<220>
<221> misc_binding
<222> 108..127
<223> 2-10-107.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 423..443
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 95..119
<223> 2-10-107 potential probe

<400> 101
caaaactgagc aaatcgcccta atcttccaaa ttctttttcc tggcttggtta gagcagccaa      60
gggtgggggtg gagcttggtga ataaaaagcc tgcttcatct tcctcaygca ggagaacatg      120
gccaagcgag ttgccattgt gggagctggg gtcagcggcc tggcctccat caagtgtgtg      180
ctggaagaag gactggagcc cacctgcttt gagaggagcg atgaccttgg ggggctgtgg      240
agattcaccg taagtgggggt ttcaacaact ttatctgtct atggagaatg gcttggcagc      300

```

87

tgggaaatta tatctgtgct tctttcacia	gggttggtgg ccttgaggaa ggtagaaat	360
gtctgtgtaa caggggacca tgaggagcca	ctgaaattgt aaaagaaaca ggacatgggt	420
gagctagggt ggaagtcaga acaggtcata		450

<210> 102  
 <211> 450  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 377  
 <223> 2-10-378 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 357..376  
 <223> 2-10-378.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 378..397  
 <223> 2-10-378.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 423..443  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 365..389  
 <223> 2-10-378 potential probe

<400> 102		
caaaactgagc aaatcgcccta atcttccaaa	ttctttttcc tggcttggtta gagcagccaa	60
gggtgggggtg gagcttgtga ataaaaagcc	tgcttcattc tcctcatgca ggagaacatg	120
gccaagcgag ttgccattgt gggagctggg	gtcagcggcc tggcctccat caagtgtgtg	180
ctggaagaag gactggagcc cacctgcttt	gagaggagcg atgaccttg ggggctgtgg	240
agattcaccg taagtggggt ttcaacaact	ttatctgtct atggagaatg gcttggcagc	300
tgggaaatta tatctgtgct tctttcacia	gggttggtgg ccttgaggaa ggtagaaat	360
gtctgtgtaa caggggrcca tgaggagcca	ctgaaattgt aaaagaaaca ggacatgggt	420
gagctagggt ggaagtcaga acaggtcata		450

<210> 103  
 <211> 446  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 284  
 <223> 2-11-284 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 264..283  
 <223> 2-11-284.mis1, potential

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<220>
<221> misc_binding
<222> 285..304
<223> 2-11-284.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 429..446
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 272..296
<223> 2-11-284 potential probe

<400> 103
tcagctagtc agactcacat ttattcggta agattaatta acagaagctt gagtcaacac      60
cgtttagagg taattgatat tatggacttc ccaagtaaaa agcacttaag cacctgccgt      120
acatcaaaagg ttagtthttaa gatcacatga gtaaacaacac taggtaggta aactactctg      180
ccttcctttg ttactacttt aatttggtta actaaaggta aagatcaggt tgccttcaac      240
catatctatt actgaagtta tgcaaacttc tcggccttcc aagragattt gtgtctatct      300
cataactat ctttaatat tttcccacca gcctgattga accccagcat agatatttaa      360
taaaaatttg gccattccgt tgttggaagt tttaagataa atttattatt attattttaa      420
tgagaaaaagc ctcaagtaaag ctgact                                     446

<210> 104
<211> 446
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 156
<223> 2-11-156 : polymorphic base A or C

<220>
<221> misc_binding
<222> 136..155
<223> 2-11-156.mis1, potential

<220>
<221> misc_binding
<222> 157..176
<223> 2-11-156.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 429..446
<223> downstream amplification primer, complement

<220>
<221> misc_binding

```

&lt;222&gt; 144..168

&lt;223&gt; 2-11-156 potential probe

&lt;400&gt; 104

tcagctagtc agactcacat ttattcggta agattaatta acagaagctt gagtcaacac	60
cgtagaggg taattgatat tatggacttc ccaagtaaaa agcacttaag cacctgccgt	120
acatcaaagg ttagttttaa gatcacatga gtaaaaaaac taggtaggta aactactctg	180
ccttcctttg ttactacttt aatttgttta actaaaggta aagatcaggt tgccttcaac	240
catatctatt actgaagtta tgcaaaacttc tcggccttcc aagaagattt gtgtctatct	300
ccataactat ctttaatat tttcccacca gcctgattga accccagcat agatatttaa	360
taaaaatttg gccattccgt tgttggaagt tttaagataa atttattatt attattttaa	420
tgagaaaagc ctcagtaaag ctgact	446

&lt;210&gt; 105

&lt;211&gt; 446

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 379

&lt;223&gt; 2-11-379 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 359..378

&lt;223&gt; 2-11-379.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 380..399

&lt;223&gt; 2-11-379.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 429..446

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 367..391

&lt;223&gt; 2-11-379 potential probe

&lt;400&gt; 105

tcagctagtc agactcacat ttattcggta agattaatta acagaagctt gagtcaacac	60
cgtagaggg taattgatat tatggacttc ccaagtaaaa agcacttaag cacctgccgt	120
acatcaaagg ttagttttaa gatcacatga gtaaaaaaac taggtaggta aactactctg	180
ccttcctttg ttactacttt aatttgttta actaaaggta aagatcaggt tgccttcaac	240
catatctatt actgaagtta tgcaaaacttc tcggccttcc aagaagattt gtgtctatct	300
ccataactat ctttaatat tttcccacca gcctgattga accccagcat agatatttaa	360
taaaaatttg gccattccrt tgttggaagt tttaagataa atttattatt attattttaa	420
tgagaaaagc ctcagtaaag ctgact	446

&lt;210&gt; 106

&lt;211&gt; 423

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

```

<220>
<221> allele
<222> 223
<223> 2-12-223 : polymorphic base A or T

<220>
<221> misc_binding
<222> 203..222
<223> 2-12-223.mis1, potential

<220>
<221> misc_binding
<222> 224..243
<223> 2-12-223.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 399..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 211..235
<223> 2-12-223 potential probe

<400> 106
ctgttcgagg tatttcttgg aaatgagcta ttacagcaa ggggtgtttgc ctctcattgc      60
tgtagttccc tgagaaaaga gcctgtgttc aatgatgagc tcccatcccg catcctgtgt      120
ggcactctgt ccatcaagcc cagtgtgaag gagtccacgg aaacctcagc tgtgtttgag      180
gatgggacca tgtttgaggc tatcgactct gtcactttg cawcaggcta tgattattcc      240
tacccttccc ttgatgagac catcatgaaa agcagaaaca atgagggtac cttgtttaa      300
ggcatcttcc cccactaat ggagaagcca accttggtg tgattggctt ggttcagtc      360
cttggagctg ccatccccc agcagacctg caagcctggt gggctgctaa agtatttgca      420
ggt                                                                423

<210> 107
<211> 418
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 239
<223> 2-14-239 : polymorphic base C or T

<220>
<221> misc_binding
<222> 219..238
<223> 2-14-239.mis1, potential

<220>
<221> misc_binding
<222> 240..259
<223> 2-14-239.mis2, potential complement

<220>
<221> primer_bind
<222> 1..23

```

```

<223> upstream amplification primer

<220>
<221> primer_bind
<222> 398..418
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 227..251
<223> 2-14-239 potential probe

<400> 107
gttcactgaa rggvacacaa ttcttggtt tctctttaag ctttcttatt ctccctagga      60
ccacacagaa gaaggcagag ccagcattta ccagtctgta ttcacaaact cttccaaaga      120
aatgatgtgc tttccagact tcccttatcc ggatgattac ccaaactata tacaccacag      180
caagctccag gaatatataa agacatatgc tcaaaagaag gatcttttaa gatacataya      240
gtttgaggta ggggtctcat aacttggtact gttgaaatta agatatgtgt gggttagaga      300
aaaaggaggc agcaaactat tataaaaatt agagccaaat gtttgggcac ctcagtaatc      360
aaatgttggc tctgattata aagcattcat gcattgattt tttctcctag acttacta      418

<210> 108
<211> 418
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 370
<223> 2-14-370 : polymorphic base G or C

<220>
<221> misc_binding
<222> 350..369
<223> 2-14-370.mis1, potential

<220>
<221> misc_binding
<222> 371..390
<223> 2-14-370.mis2, potential complement

<220>
<221> primer_bind
<222> 1..23
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 398..418
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 358..382
<223> 2-14-370 potential probe

<400> 108
gttcactgaa rggvacacaa ttcttggtt tctctttaag ctttcttatt ctccctagga      60
ccacacagaa gaaggcagag ccagcattta ccagtctgta ttcacaaact cttccaaaga      120
aatgatgtgc tttccagact tcccttatcc ggatgattac ccaaactata tacaccacag      180
caagctccag gaatatataa agacatatgc tcaaaagaag gatcttttaa gatacataya      240
gtttgaggta ggggtctcat aacttggtact gttgaaatta agatatgtgt gggttagaga      300
aaaaggaggc agcaaactat tataaaaatt agagccaaat gtttgggcac ctcagtaatc      360

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92

aaatgttggs tctgattata aagcattcat gcattgattt tttctcctag acttacta 418

&lt;210&gt; 109

&lt;211&gt; 445

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 104

&lt;223&gt; 2-17-104 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 84..103

&lt;223&gt; 2-17-104.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 105..124

&lt;223&gt; 2-17-104.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..18

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 427..445

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 92..116

&lt;223&gt; 2-17-104 potential probe

&lt;400&gt; 109

cggtgatgga aaattcccct ctgctctctg aagatttgct aaaaatctac tgacaaaagg	60
catattgata gaagaaaatg tacacaaatt tattaacttc cacrggagtt ggggtaaaaa	120
tcacatgatt atcccagcat gcaatggggg acggatgctt atatatccct tccttaggtg	180
acagggagat gagaaagtgt ggattgattt tagggtgact atgaaatgat ctctagggga	240
cccaacgggc ttgaagaaca tacaatggcc tggaataaag tatgttgggc ccgcagcgca	300
aacaatggct tatgacaagt ctgtctaggt gtgttgacag aattctttct tcctgcagta	360
tgagttcagt taatgaaaac tcaggggaagg taccaaagggt aattgatttc ttctttggca	420
agtctagact ttaggcaaat aagg	445

&lt;210&gt; 110

&lt;211&gt; 445

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 395

&lt;223&gt; 2-17-396 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 375..394

&lt;223&gt; 2-17-396.mis1, potential

&lt;220&gt;



```

<221> misc_binding
<222> 396..415
<223> 2-17-396.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 427..445
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 383..407
<223> 2-17-396 potential probe

<400> 110
cgttgatgga aaattcccct ctgctctctg aagatttgct aaaaatctac tgacaaaagg      60
catattgata gaagaaaatg tacacaaatt tattaacttc cacaggagtt ggggtaaaaa      120
tcacatgatt atcccagcat gcaatggggt acggatgctt atatatccct tccttaggtg      180
acagggagat gagaaagtgt ggattgattt tagggtgact atgaaatgat ctctagggga      240
cccaacgggc ttgaagaaca tacaatggcc tggataaaag tatgttgggc ccgcagcgca      300
aacaatggct tatgacaagt ctgtctaggt gtgttgacag aattctttct tcctgcagta      360
tgagttcagt taatgaaaac tcaggaaggg taccmaaggt aattgatttc ttctttggca      420
agtctagact ttaggcaaat aagggg                                     445

<210> 111
<211> 436
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 43
<223> 2-22-43 : polymorphic base A or G

<220>
<221> misc_binding
<222> 23..42
<223> 2-22-43.mis1, potential

<220>
<221> misc_binding
<222> 44..63
<223> 2-22-43.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 416..436
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 31..55
<223> 2-22-43 potential probe

```

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<400> 111
aaggagttca cggaaacctc agctgtgttt gaggatggga ccrtgtttga ggctatcgac    60
tctgtcatct ttgcaacagg ctatgattat tcctaccctt tccttgatga gaccatcatg    120
aaaagcagaa acaatgaggt taccttggtt aaaggcatct tccccccact aatggagaag    180
ccaaccttgg ctgtgattgg cttggttcag tcccttggag ctgccatccc cacagcagac    240
ctgcaagcct ggtgggctgc taaagtattt gcaagtaggt gggccattct gtctttcatt    300
cattttatca atgaacattt actgaacacc tgctatatgc aaagcactgt gctagggata    360
caatgagaac aagacaaaca tggtccttga cctctcaagg cttaaaatgg ggtgtggggg    420
atgcataata ggggaa                                     436

```

```

<210> 112
<211> 436
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele
<222> 138
<223> 2-22-138 : polymorphic base A or G

```

```

<220>
<221> misc_binding
<222> 118..137
<223> 2-22-138.mis1, potential

```

```

<220>
<221> misc_binding
<222> 139..158
<223> 2-22-138.mis2, potential complement

```

```

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

```

```

<220>
<221> primer_bind
<222> 416..436
<223> downstream amplification primer, complement

```

```

<220>
<221> misc_binding
<222> 126..150
<223> 2-22-138 potential probe

```

```

<400> 112
aaggagttca cggaaacctc agctgtgttt gaggatggga ccattgtttga ggctatcgac    60
tctgtcatct ttgcaacagg ctatgattat tcctaccctt tccttgatga gaccatcatg    120
aaaagcagaa acaatgargt taccttggtt aaaggcatct tccccccact aatggagaag    180
ccaaccttgg ctgtgattgg cttggttcag tcccttggag ctgccatccc cacagcagac    240
ctgcaagcct ggtgggctgc taaagtattt gcaagtaggt gggccattct gtctttcatt    300
cattttatca atgaacattt actgaacacc tgctatatgc aaagcactgt gctagggata    360
caatgagaac aagacaaaca tggtccttga cctctcaagg cttaaaatgg ggtgtggggg    420
atgcataata ggggaa                                     436

```

```

<210> 113
<211> 420
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele

```

```

<222> 82
<223> 2-23-82 : polymorphic base A or G

<220>
<221> misc_binding
<222> 62..81
<223> 2-23-82.mis1, potential

<220>
<221> misc_binding
<222> 83..102
<223> 2-23-82.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 70..94
<223> 2-23-82 potential probe

<400> 113
caattatttc ctccacagaa acaaggcaag aaggaaaaaa actttcacat gtagaattat      60
aaatggaaaa ataaattttc trgttttctt aaagaccctg gtttccggta taaagaaatg      120
tcccagcttc ttagtcacgg gccaatgggt tgttgttact gaaaaggatg ggaaacagga      180
atctactatt tttgatgctg taatgatttg ttcaggacat cacgtatacc ccaatctgcc      240
aacggattcc tttcctggta agtttggaat atatataata atctagggac ttatatgcaa      300
acatcaagag ttagaaacat atctttctat aggtattaca taatgattat tcttagattt      360
caaaagaaaa aaattaagtt taatgatagg atatagtaat aaatagccyc ataagtcctt      420

<210> 114
<211> 420
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 166
<223> 2-23-166 : polymorphic base A or G

<220>
<221> misc_binding
<222> 146..165
<223> 2-23-166.mis1, potential

<220>
<221> misc_binding
<222> 167..186
<223> 2-23-166.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>

```

96

```

<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 154..178
<223> 2-23-166 potential probe

<400> 114
caattatttc ctccacagaa acaaggcaag aaggaaaaaa actttcacat gtagaattat    60
aaatggaaaa ataaattttc tagttttctt aaagaccctg gtttcggtg taaagaaatg    120
tcccagcttc ttagtcacgg gccaatgggt tgttggtact gaaaargatg ggaaacagga    180
atctactatt ttgatgctg taatgatttg ttcaggacat cacgtatacc ccaatctgcc    240
aacggattcc tttcctggta agtttggaat atatataata atctagggac ttatatgcaa    300
acatcaagag ttagaaacat atctttctat aggtattaca taatgattat tcttagattt    360
caaaagaaaa aaattaagtt taatgatagg atatagtaat aaatagccyc ataagtcctt    420

<210> 115
<211> 420
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 244
<223> 2-23-244 : polymorphic base G or T

<220>
<221> misc_binding
<222> 224..243
<223> 2-23-244.mis1, potential

<220>
<221> misc_binding
<222> 245..264
<223> 2-23-244.mis2, potential complement

<220>
<221> primer_bind
<222> 1..25
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 232..256
<223> 2-23-244 potential probe

<400> 115
caattatttc ctccacagaa acaaggcaag aaggaaaaaa actttcacat gtagaattat    60
aaatggaaaa ataaattttc tagttttctt aaagaccctg gtttcggtg taaagaaatg    120
tcccagcttc ttagtcacgg gccaatgggt tgttggtact gaaaaggatg ggaaacagga    180
atctactatt ttgatgctg taatgatttg ttcaggacat cacgtatacc ccaatctgcc    240
aackgattcc tttcctggta agtttggaat atatataata atctagggac ttatatgcaa    300
acatcaagag ttagaaacat atctttctat aggtattaca taatgattat tcttagattt    360
caaaagaaaa aaattaagtt taatgatagg atatagtaat aaatagccyc ataagtcctt    420

<210> 116

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<211> 434
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 115
<223> 2-24-115 : polymorphic base C or T

<220>
<221> misc_binding
<222> 95..114
<223> 2-24-115.mis1, potential

<220>
<221> misc_binding
<222> 116..135
<223> 2-24-115.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 416..434
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 103..127
<223> 2-24-115 potential probe

<400> 116
aatgcacttc aactagggaa ttttttaatt acaactgata ataggtttaa aaagacacaa      60
agaaaacatc ttcataatct ctgaaaatca gttcaaacaa cttgccatgt tccayttagg      120
cctggaccag ttctgaggca actacctcca tagccgggat tataagaatc cagaagcctt      180
caaggggaag agggctcctcg tgattggtct ggggaattcg ggatctgaca ttgctgttga      240
gctcagccgt ctggctacac aggtacatga cgtaaagggt ttgggaaata aacctaagggt      300
agggctgtgc tactaaatca gtagccaagg cacagaggat ggtacttcta tgtcacacca      360
caagagatcc acctcttcta tgtggccctt caaatcaagg aggacttgag acatcctcca      420
tgtgaagcca ggta                                         434

<210> 117
<211> 420
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 36
<223> 2-25-36 : polymorphic base G or C

<220>
<221> misc_binding
<222> 16..35
<223> 2-25-36.mis1, potential

<220>
<221> misc_binding
<222> 37..56
<223> 2-25-36.mis2, potential complement

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<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 396..420
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 24..48
<223> 2-25-36 potential probe

<400> 117
aggaaaatgc acggaagtgc ccaaagaagt gtgttstggt tgettatttc ttacagagta      60
atgctgaaat ctgtgttgct tttcccacc aggtcattat cagtaccaga agtgcttcct      120
gggtcatgag tcgggtctgg gatgatggct atccttggga tatgatgtat gttaccgcgt      180
ttgcatcctt tctccggaat gtccttcctt cattcatctc tgactgggta tatgtccaga      240
agatgaacac gtggtttaag catgagaact atggcctgat gcctttaaat gggactactaa      300
aaatggaaat tttttttatt caaaaaaggg gggcactcat ttaatgaatt tattctctct      360
agaacttact tttgttgtct cattgagcct agaaacatta aactcaaggt ttcacagggt      420

<210> 118
<211> 429
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 185
<223> 2-27-185 : polymorphic base A or G

<220>
<221> misc_binding
<222> 165..184
<223> 2-27-185.mis1, potential

<220>
<221> misc_binding
<222> 186..205
<223> 2-27-185.mis2, potential complement

<220>
<221> primer_bind
<222> 1..19
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 405..429
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 173..197
<223> 2-27-185 potential probe

<400> 118
cttattccag tatgttctct tctttcttca tgtttggcca gagccagact ttgcagacag      60
attacatcac atatgtggat gagctgggct ctttcatagg ggccaagcct aacataccat      120

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99

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ggctcttctt gacagatccc cgcttggccc tggaggtgta ctttggccct tgcagcccat 180
accarttttcg actgatggga ccaggaaggt gggatggggc cagaaatgcc atcctgacct 240
agtgggaaccg gacagtgaag ccaaccagga caagagttgt cagtgaagtt cagcgacccc 300
atcccttttta caatttgctt aaaatgcttt cattcccatt actccttctg gctgttacac 360
ttacatttta ttaatgagaa agtctttgag gtctcaaaat tcagcataga agtgtaatca 420
cacaatata 429

```

&lt;210&gt; 119

&lt;211&gt; 429

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 378

&lt;223&gt; 2-27-378 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 358..377

&lt;223&gt; 2-27-378.misl, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 379..398

&lt;223&gt; 2-27-378.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..19

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 405..429

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 366..390

&lt;223&gt; 2-27-378 potential probe

&lt;400&gt; 119

```

cttattccag tatgttctct tctttcttca tgtttggcca gagccagact ttgcagacag 60
attacatcac atatgtggat gagctgggct ctttcatagg ggccaagcct aacataccat 120
ggctcttctt gacagatccc cgcttggccc tggaggtgta ctttggccct tgcagcccat 180
accagttttcg actgatggga ccaggaaggt gggatggggc cagaaatgcc atcctgacct 240
agtgggaaccg gacagtgaag ccaaccagga caagagttgt cagtgaagtt cagcgacccc 300
atcccttttta caatttgctt aaaatgcttt cattcccatt actccttctg gctgttacac 360
ttacatttta ttaatgasaa agtctttgag gtctcaaaat tcagcataga agtgtaatca 420
oacaatata 429

```

&lt;210&gt; 120

&lt;211&gt; 440

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 142

&lt;223&gt; 2-29-142 : polymorphic base A or G

&lt;220&gt;

100

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<221> misc_binding
<222> 122..141
<223> 2-29-142.mis1, potential

<220>
<221> misc_binding
<222> 143..162
<223> 2-29-142.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 422..439
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 130..154
<223> 2-29-142 potential probe

<220>
<221> misc_feature
<222> 9,433
<223> n=a, g, c or t

<400> 120
caattawtna ctccagaaag gaaaagctgg caatgcagtt ttattgaaat tagcttgaca      60
tagttgctct ggagctcaca gacttctctc ttcttcccc tgaaggatat gagagggttca      120
aaggccaata tttccatagc crccaataca agcatccaga tggatttggg gaaaacgcat      180
cctgggtgatt ggaatgggaa acttgggctc agatattgct gttgagctga gtaagaatgc      240
tgctcaggtg tgatgctctc tgcttaccat gtacctggag gggaggaagt ggggatgcca      300
tactggagaa ccycagccat ataatcgcgg ctccaatcct cattaactag ttgggttgga      360
gcgcattgtg gcatcataga aaatctggaa gtcaagaaac cactttacct cctagctctg      420
tcactaacca gcnatgaatg                                     440

<210> 121
<211> 440
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 166
<223> 2-29-166 : polymorphic base C or T

<220>
<221> misc_binding
<222> 146..165
<223> 2-29-166.mis1, potential

<220>
<221> misc_binding
<222> 167..186
<223> 2-29-166.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

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<220>
<221> primer_bind
<222> 422..439
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 154..178
<223> 2-29-166 potential probe

<220>
<221> misc_feature
<222> 9,433
<223> n=a, g, c or t

<400> 121
caattawtna ctccagaaag gaaaagctgg caatgcagtt ttattgaaat tagcttgaca      60
tagttgctct ggagctcaca gacttctctc ttcttcccc tgaaggatat gagaggttca      120
aaggccaata ttccatagc cgccaataca agcatccaga tggatytggg gaaaacgcat      180
cctggtgatt ggaatgggaa acttgggctc agatattgct gttgagctga gtaagaatgc      240
tgctcagggtg tgatgctctc tgcttaccat gtacctggag gggaggaagt ggggatgcca      300
tactggagaa ccycagccat ataatcgcggt ctccaatcct cattaactag ttggttggtgta      360
gcgcattgtg gcatcataga aaatctggaa gtcaagaaac cactttacct cctagctctg      420
tcactaacca gcnatgaatg                                     440

<210> 122
<211> 440
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 204
<223> 2-29-205 : polymorphic base C or T

<220>
<221> misc_binding
<222> 184..203
<223> 2-29-205.mis1, potential

<220>
<221> misc_binding
<222> 205..224
<223> 2-29-205.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 422..439
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 192..216
<223> 2-29-205 potential probe

<220>
<221> misc_feature

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102

&lt;222&gt; 9,433

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 122

caattawtna ctccagaaag gaaaagctgg caatgcagtt ttattgaaat tagcttgaca	60
tagttgctct ggagctcaca gacttctctc ttcttcccc tgaaggatat gagaggttca	120
aaggccaata tttccatagc cgccaatata agcatccaga tggatttggg gaaaacgcat	180
cctggtgatt ggaatgggaa actygggctc agatattgct gttgagctga gtaagaatgc	240
tgctcagggtg tgatgctctc tgcttaccat gtacctggag gggaggaaagt ggggatgcca	300
tactggagaa ccycagccat ataatcgagg ctccaatcct cattaactag ttggttggtg	360
gcgcattgtg gcatcataga aaatctggaa gtcaagaaac cactttacct cctagctctg	420
tcactaacca gcnatgaatg	440

&lt;210&gt; 123

&lt;211&gt; 440

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 205

&lt;223&gt; 2-29-206 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 185..204

&lt;223&gt; 2-29-206.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 206..225

&lt;223&gt; 2-29-206.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..21

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 422..439

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 193..217

&lt;223&gt; 2-29-206 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 9,433

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 123

caattawtna ctccagaaag gaaaagctgg caatgcagtt ttattgaaat tagcttgaca	60
tagttgctct ggagctcaca gacttctctc ttcttcccc tgaaggatat gagaggttca	120
aaggccaata tttccatagc cgccaatata agcatccaga tggatttggg gaaaacgcat	180
cctggtgatt ggaatgggaa acttrggctc agatattgct gttgagctga gtaagaatgc	240
tgctcagggtg tgatgctctc tgcttaccat gtacctggag gggaggaaagt ggggatgcca	300
tactggagaa ccycagccat ataatcgagg ctccaatcct cattaactag ttggttggtg	360
gcgcattgtg gcatcataga aaatctggaa gtcaagaaac cactttacct cctagctctg	420
tcactaacca gcnatgaatg	440

103

<210> 124  
 <211> 440  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 313  
 <223> 2-29-314 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 293..312  
 <223> 2-29-314.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 314..333  
 <223> 2-29-314.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..21  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 422..439  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 301..325  
 <223> 2-29-314 potential probe

<220>  
 <221> misc\_feature  
 <222> 9,433  
 <223> n=a, g, c or t

<400> 124	
caattawtna ctccagaaag gaaaagctgg caatgcagtt ttattgaaat tagcttgaca	60
tagttgctct ggagctcaca gacttctctc ttcttcccc tgaaggtagt gagaggttca	120
aaggccaata tttccatagc cgccaataca agcatccaga tggatttggg gaaaacgcat	180
cctgggtgatt ggaatgggaa acttgggctc agatattgct gttgagctga gtaagaatgc	240
tgctcaggtg tgatgctctc tgcttaccat gtacctggag gggaggaagt ggggatgcca	300
tactggagaa ccycagccat ataatcgcg ctccaatcct cattaactag ttggttggtg	360
gcgcattgtg gcatcataga aaatctggaa gtcaagaaac cactttacct cctagctctg	420
tcactaacca gcnatgaatg	440

<210> 125  
 <211> 432  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 68  
 <223> 2-32-68 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 48..67

<223> 2-32-68.mis1, potential

<220>

<221> misc\_binding

<222> 69..88

<223> 2-32-68.mis2, potential complement

<220>

<221> primer\_bind

<222> 1..21

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 413..432

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 56..80

<223> 2-32-68 potential probe

<400> 125

cctatgagta	tcgcctgggt	gggcctgggc	aatgggaagg	agccagaaat	gccatcttca	60
cccagaarca	aagaatactg	aagccactca	agactcgggc	cctgaaggat	tcatactaatt	120
tctcagtttc	ttttctgttg	aaaatcctgg	gccttccttg	tggtgtgtg	gccttttttt	180
gccaaacttca	atggtcctag	tcagcataat	gctttgggct	ttattatctt	gtcagtcact	240
acctcctaaa	gaaaaaaaaa	aaggctagaa	gaaaaaacat	tacattcatg	ttctaattat	300
agatttttaga	gttaggtagt	acagghaagg	gggaaattgt	aaagaattag	cagaattagg	360
catatgtaca	aaaccaaata	tttgtcatga	aattttgcct	ttccacgctt	ccctcagttc	420
accaaagtta	cc					432

<210> 126

<211> 424

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 357

<223> 2-35-357 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 337..356

<223> 2-35-357.mis1, potential

<220>

<221> misc\_binding

<222> 358..377

<223> 2-35-357.mis2, potential complement

<220>

<221> primer\_bind

<222> 1..18

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 404..423

<223> downstream amplification primer, complement

<220>

105

&lt;221&gt; misc\_binding

&lt;222&gt; 345..369

&lt;223&gt; 2-35-357 potential probe

&lt;400&gt; 126

gaagagccta ttgacatcat tgtctttgcc actggataca catttgcttt ccccttcctt	60
gatgagtctg tagtgaaagt tgaagatggc caggcctcac tgtacaagta tatcttcctt	120
gcacatctgc aaaagccaac cctggccatt attggcctca tcaaaccctt gggctccatg	180
atacctacag gagaaacaca agctcgggtgg gctgttcgag tcctgaaagg taagtataag	240
aaatagcagg gcatgtgttt ttggtgtgcc atgtgattct ggatactgga aatgttgaga	300
ctattattcc tcctgcttct atttaaaata acagaatctt taaaagcagg atgcatycta	360
ttgtttgctg aattatactg tcataatgat ttgttcatt actgcataaa tggatatatca	420
gggg	424

&lt;210&gt; 127

&lt;211&gt; 435

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 256

&lt;223&gt; 2-36-256 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 236..255

&lt;223&gt; 2-36-256.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 257..276

&lt;223&gt; 2-36-256.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..21

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 411..435

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 244..268

&lt;223&gt; 2-36-256 potential probe

&lt;400&gt; 127

atatcaatgg tacatcaact ggcaagtgct taatggaaca ttgacaactg ctaatcatat	60
ctgtgattct tttttcagac caaagtctgc agtgtaacaa aatgctcaga ttctgctgtc	120
tctggccaat gggagggtgg cactatgcat gaagagaagc aagagtcagc catctttgat	180
gctgtcatgg tctgcactgg ctttcttact aatccttatt tgccactgga ttcctttcca	240
ggtacagcat tttctstaac taactttaag ttttctcgtg ggagccattc tgatgcttga	300
ttggtctggg aatgaattcc tatggctgtt ccattaaata gttaaagttg ggaggttagga	360
ggaggctttt ttgtttttgt ttgtttttt tctagccagc attttctggc cagtttttgg	420
ctttcatttg ttcca	435

&lt;210&gt; 128

&lt;211&gt; 435

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

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<220>
<221> allele
<222> 353
<223> 2-36-354 : polymorphic base A or C

<220>
<221> misc_binding
<222> 333..352
<223> 2-36-354.mis1, potential

<220>
<221> misc_binding
<222> 354..373
<223> 2-36-354.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 411..435
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 341..365
<223> 2-36-354 potential probe

<400> 128
atatcaatgg tacatcaact ggcaagtgtc taatggaaca ttgacaactg ctaatcatat      60
ctgtgattct tttttcagac caaagtctgc agtgtaacaa aatgctcaga ttctgctgtc      120
tctggccaat gggaggtggt cactatgcat gaagagaagc aagagtcagc catctttgat      180
gctgtcatgg tctgcactgg ctttcttact aatccttatt tgccactgga ttcctttcca      240
ggtacagcat tttctgtaac taactttaag ttttctcgtg ggagccattc tgatgcttga      300
ttggtctggg aatgaattcc tatggctgtt ccattaaata gttaaagttg ggmggtagga      360
ggaggctttt ttgttttgt tttgtttttt tctagccagc attttctggc cagtttttgg      420
ctttcatttg ttcca                                     435

<210> 129
<211> 449
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 236
<223> 2-42-236 : polymorphic base C or T

<220>
<221> misc_binding
<222> 216..235
<223> 2-42-236.mis1, potential

<220>
<221> misc_binding
<222> 237..256
<223> 2-42-236.mis2, potential complement

<220>
<221> primer_bind

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<222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 428..449  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 224..248  
 <223> 2-42-236 potential probe

<400> 129  
 agtggggtttg cgtggttggg ggcagatttt ttccccact cacacttttg gcaactsacag 60  
 tttttcagct gtctcatgga gtttgcagcg gcaagccact tctttcaaag gtcctgtgaa 120  
 ttcttttggg tttcctggta tgttcagca gtacttcttg gaggaaaatt tcaactgtgtg 180  
 agtctccaca tgctgttctg tctgtctgcg tggaaactgc aagttagtcc tgccctyctat 240  
 ccaccatttt ccaacaatct gtcgttaacc atttcaaagt atagaattca gtggcattta 300  
 gtacattcat aatgctgtgt aaccacaacc tctatccagt ttcaaaacac ttccatcaca 360  
 cccaaaagaa aactccatac tcattagcaa tccttcccca ttcccttctt tccccagccc 420  
 ctggcagcca ctgatcatg cctatttta 449

<210> 130  
 <211> 429  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 139  
 <223> 2-43-139 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 119..138  
 <223> 2-43-139.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 140..159  
 <223> 2-43-139.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 395..419  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 127..151  
 <223> 2-43-139 potential probe

<400> 130  
 cattakcaat ccttccccat tcccttcttt cccagcccc tggcagccac tcgatcatgc 60  
 ctattttaaa ttgaggaaac taaagctgag aaaagttata caatttttcc aacatgactc 120  
 tgataatagc tgggtgaacrc aatactcgaa cccaggactt gtgattccca agatcagact 180  
 cctcccatat acctgtcttt tatttcccaa gtgcttgggc aatcagctgc acttaaaaag 240

108

cctttacaaa	ttgagtttac	taatgtgagt	aatatatgat	tttttaaaaa	taataattgt	300
ccctaaaggt	gaaatggatc	aaagccctta	aaagtgaatc	tgtgggtgtag	taactgttaa	360
cataattgtc	ttatttttatt	ctcatcccct	taaagaataa	aattgatgaa	caagtgaggg	420
gtcatagct						429

<210> 131  
 <211> 425  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 213  
 <223> 2-44-215 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 193..212  
 <223> 2-44-215.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 214..233  
 <223> 2-44-215.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 3..23  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 398..420  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 201..225  
 <223> 2-44-215 potential probe

<400> 131							
rhcctgcttt	tatttcccaa	gtgcttgggc	aatcagctgc	acttaaaaag	cctttacaaa		60
ttgagtttac	taatgtgagt	aatatatgat	tttttaaaaa	taataattgt	ccctaaaggt		120
gaaatggatc	aaagccctta	aaagtgaatc	tgtgggtgtag	taactgttaa	cataattgtc		180
ttatttttatt	ctcatcccct	taaagaataa	aatgtgatgaa	caagtgagga	tgctgggtgta		240
actccctaac	ttagttttata	gtctgtaagc	agaagagtga	gtctaaagta	catatcacca		300
gacagtgttt	ctcctagcat	ttcgctgttg	cattaatcaa	caagttaaaa	tataaacaac		360
ggctaaccct	gggtttcaaa	tttaacattc	cttatctctt	agaccagggt	tatccactgt		420
ggtca							425

<210> 132  
 <211> 442  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 38  
 <223> 2-45-38 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 18..37



109

&lt;223&gt; 2-45-38.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 39..58

&lt;223&gt; 2-45-38.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..21

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 418..442

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 26..50

&lt;223&gt; 2-45-38 potential probe

<400> 132	
gatgtaggaa gagtaaaaaa caaaaaattt ttgaatgygt aattatcact aattatttta	60
tttgatcctt caggagaatg tggaagatgg ccgagcaagt atctatcaat ctgtcgttac	120
caacaccagc aaagaaatgt cctgtttcag tgactttcca atgcctgaag attttccaaa	180
cttcctgcat aattctaaac ttctggaata ttccaggatt ttgctaaaa aatttgatct	240
gctaaaatat attcagttcc aggtattgta tttttgggga aatgggtttc tctgcattag	300
ttcagctcat atttagatag aaaagttact ctgataatga aagcaattat gaatgaagta	360
tcccattcta agtattttgt gaaatataac agcctcatat aaaacccaaa aagtagtgtc	420
attacccttg gtattataga tt	442

&lt;210&gt; 133

&lt;211&gt; 442

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 183

&lt;223&gt; 2-45-183 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 163..182

&lt;223&gt; 2-45-183.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 184..203

&lt;223&gt; 2-45-183.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..21

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 418..442

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

110

&lt;221&gt; misc\_binding

&lt;222&gt; 171..195

&lt;223&gt; 2-45-183 potential probe

&lt;400&gt; 133

gatgtaggaa gagtaaaaaa caaaaaat	ttgaatgcgt aattatcact aattatttta	60
tttgatcctt caggagaatg tggaagatgg	ccgagcaagt atctatcaat ctgtcggttac	120
caacaccagc aaagaaatgt cctgtttcag	tgactttcca atgcctgaag attttccaaa	180
ctycctgcat aattctaaac ttctggaata	tttcaggatt ttgctaaaa aatttgatct	240
gctaaaatat attcagttcc aggtattgta	tttttgggga aatgggtttc tctgcattag	300
ttcagctcat atttagatag aaaagttact	ctgataatga aagcaattat gaatgaagta	360
tcccattcta agtatttggt gaaatataac	agcctcatat aaaacccaaa aagtagtgtc	420
attacccttg gtattataga tt		442

&lt;210&gt; 134

&lt;211&gt; 442

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 335

&lt;223&gt; 2-45-335 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 315..334

&lt;223&gt; 2-45-335.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 336..355

&lt;223&gt; 2-45-335.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..21

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 418..442

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 323..347

&lt;223&gt; 2-45-335 potential probe

&lt;400&gt; 134

gatgtaggaa gagtaaaaaa caaaaaat	ttgaatgcgt aattatcact aattatttta	60
tttgatcctt caggagaatg tggaagatgg	ccgagcaagt atctatcaat ctgtcggttac	120
caacaccagc aaagaaatgt cctgtttcag	tgactttcca atgcctgaag attttccaaa	180
cttcctgcat aattctaaac ttctggaata	tttcaggatt ttgctaaaa aatttgatct	240
gctaaaatat attcagttcc aggtattgta	tttttgggga aatgggtttc tctgcattag	300
ttcagctcat atttagatag aaaagttact	ctgawaatga aagcaattat gaatgaagta	360
tcccattcta agtatttggt gaaatataac	agcctcatat aaaacccaaa aagtagtgtc	420
attacccttg gtattataga tt		442

&lt;210&gt; 135

&lt;211&gt; 442

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

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<220>
<221> allele
<222> 394
<223> 2-45-394 : polymorphic base C or T

<220>
<221> misc_binding
<222> 374..393
<223> 2-45-394.mis1, potential

<220>
<221> misc_binding
<222> 395..414
<223> 2-45-394.mis2, potential complement

<220>
<221> primer_bind
<222> 1..21
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 418..442
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 382..406
<223> 2-45-394 potential probe

<400> 135
gatgtaggaa gagtaaaaaa caaaaaatTT ttgaatgcgt aattatcact aattatttta      60
tttgatcctt caggagaatg tggaagatgg ccgagcaagt atctatcaat ctgtcggttac      120
caacaccagc aaagaaatgt cctgtttcag tgactttcca atgcctgaag attttccaaa      180
cttcctgcat aattctaaac ttctggaata tttcaggatt tttgctaaaa aatttgatct      240
gctaaaatat attcagttcc aggtattgta tttttgggga aatgggtttc tctgcattag      300
ttcagctcat atttagatag aaaagttact ctgataatga aagcaattat gaatgaagta      360
tcccattcta agtatttggt gaaatataac agcytcatat aaaacccaaa aagtagtgc      420
attacccttg gtattataga tt                                         442

<210> 136
<211> 426
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 39
<223> 2-48-39 : polymorphic base A or G

<220>
<221> misc_binding
<222> 19..38
<223> 2-48-39.mis1, potential

<220>
<221> misc_binding
<222> 40..59
<223> 2-48-39.mis2, potential complement

<220>
<221> primer_bind

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<222> 1..24  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 404..426  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 27..51  
 <223> 2-48-39 potential probe

```

<400> 136
gttttatttt attgatgggc tgtctggctc cctcaactrc aaagtaaact ccacaaaggc      60
agagagtttt gcctctttta ttcattgctg tacctgcatc acttagaaaag tttctggcac      120
ctaggaagtg ttcagtaaat atttattgaa taagtttatg taaaacgtct cagactcctt      180
agagaaactg gtcttttggg gttggagaat aaagttcttt acctcatcag ttagactcta      240
tctaaggtac acgagggctt gctagtctcc taagttagtc tgctaataaa tgtaaccct      300
aataactgaa attattagca gaggtaatta tccagttcta tatcaaggca aaaagacagc      360
agtggataga aagatccttag aagtcccact aggttcaccc aagccaccat acacataggc      420
agaaaaa                                         426
  
```

<210> 137  
 <211> 426  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 72  
 <223> 2-48-72 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 52..71  
 <223> 2-48-72.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 73..92  
 <223> 2-48-72.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..24  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 404..426  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 60..84  
 <223> 2-48-72 potential probe

```

<400> 137
gttttatttt attgatgggc tgtctggctc cctcaactac aaagtaaact ccacaaaggc      60
agagagtttt gycctcttta ttcattgctg tacctgcatc acttagaaaag tttctggcac      120
ctaggaagtg ttcagtaaat atttattgaa taagtttatg taaaacgtct cagactcctt      180
agagaaactg gtcttttggg gttggagaat aaagttcttt acctcatcag ttagactcta      240
  
```

113

tctaaggtac acgagggcctt gctagtctcc taagttagtc tgctaataaa tgtaaccct	300
aataactgaa attattagca gaggttaatta tccagttcta tatcaaggca aaaagacagc	360
agtggataga aagatcttag aagtcacct aggttcatcc aagccacccat acacataggc	420
agaaaa	426

&lt;210&gt; 138

&lt;211&gt; 426

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 156

&lt;223&gt; 2-48-156 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 136..155

&lt;223&gt; 2-48-156.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 157..176

&lt;223&gt; 2-48-156.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..24

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 404..426

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 144..168

&lt;223&gt; 2-48-156 potential probe

&lt;400&gt; 138

gttttatttt attgatgggc tgtctggctc cctcaactac aaagtaaact ccacaaaggc	60
agagagtttt gcccttttta ttcattgctg tacctgcac accatgaaag tttctggcac	120
ctaggaagtg ttcagtaaat atttattgaa taagtktatg taaaacgtct cagactcctt	180
agagaaactg gtcttttggg gttggagaat aaagttcttt acctcatcag ttagactcta	240
tctaaggtac acgagggcctt gctagtctcc taagttagtc tgctaataaa tgtaaccct	300
aataactgaa attattagca gaggttaatta tccagttcta tatcaaggca aaaagacagc	360
agtggataga aagatcttag aagtcacct aggttcatcc aagccacccat acacataggc	420
agaaaa	426

&lt;210&gt; 139

&lt;211&gt; 426

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 285

&lt;223&gt; 2-48-285 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 265..284

114

&lt;223&gt; 2-48-285.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 286..305

&lt;223&gt; 2-48-285.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..24

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 404..426

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 273..297

&lt;223&gt; 2-48-285 potential probe

&lt;400&gt; 139

gttttatttt attgatgggc tgtctggctc cctcaactac aaagtaaact ccacaaaggc	60
agagagtttt gcctctttta ttcattgctg tacctgcatc acttagaaag tttctggcac	120
ctaggaagtg ttcagtaaat atttattgaa taagtttatg taaaacgtct cagactcctt	180
agagaaactg gtcttttggg gttggagaat aaagttcttt acctcatcag ttagactcta	240
tctaaggtag acgagggtt gctagtctcc taagttagtc tgctrataaa tgtaaccct	300
aataactgaa attattagca gaggtaatta tccagttcta tatcaaggca aaaagacagc	360
agtggataga aagatcttag aagtccact aggttcatcc aagccaccat acacataggc	420
agaaaa	426

&lt;210&gt; 140

&lt;211&gt; 420

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 167

&lt;223&gt; 2-49-167 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 147..166

&lt;223&gt; 2-49-167.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 168..187

&lt;223&gt; 2-49-167.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 1..20

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 396..420

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

115

<221> misc\_binding  
 <222> 155..179  
 <223> 2-49-167 potential probe

<400> 140  
 aagtcctcact aggttcatcc aagccacccat acacataggc agaaaaatca aaataagata 60  
 tgagcctgga caggggtgagc aatctgggaa aagatgaaca cagtatgcta ggacccagaa 120  
 atcatcaagt ctatgaaaac taagccagaa cacaaatgtg aattccrtaa gatcaggaac 180  
 ataattctgtc ttgttcatcc aggcattggta atctgccaga aatagtgtt aactgcaaga 240  
 actgaatatt tgtagataa ttaaaccatc aactaaatga gattcatgca accatgaaaa 300  
 atgctgctat aggtacacaa tattgatata ctagaaagtt aaaaaatcaa gttggaaatt 360  
 agactattcc atttctgttt gtgtgtatgt atctacaaat aggtggaakg atataccaaa 420

<210> 141  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-436-43 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 10-436-43.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-436-43.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 459..476  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 859..878  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-436-43 potential probe

<220>  
 <221> misc\_feature  
 <222> 297,539,629,650,976..1001  
 <223> n=a, g, c or t

<400> 141  
 tatatttaat acatatatatt aataaaaaaca ttaaatacat atattttaata aaaatataat 60  
 ttatatataa tatacatata tttatatata aatatattta tatataatat gcatatgttt 120  
 attttacata tataaatata tttatatata aatatacgta tattttatttt atatataat 180  
 gtgtgtgtgt atatatatat atacacacac acacgtatat acccagggag aggcactaat 240  
 tgtgcagttt tgaaaagttt cctagtgtat ctgccgtggc ctactttcaa agcactntgc 300  
 ttaggtttac cacttaaaat gttattacat tttccagaa gtacacttta aaatactttg 360  
 tttttaagt gaggcattac aatggtgtca tgagctgaca ttcccagcca ctgactggaa 420  
 gatttgggtc aaccagggga acaggcttgg ttttaaggag ttcaagtgtt ggtggccttt 480  
 gggtaaaagt ttatgtctat stttgtatgt gggattcagt agggattttc tttaatgtnt 540

116

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aaaaattcta gtttcctctg tatttgcttt ccttcttccc ctgtcctccc tgagcctcca    600
cacatgtgta acacatttta ttgccaant aggtcaagga ggtaaaaaan gactttgtcg    660
ggcttggaag tcagcatggt tactgcagtt cccggtagge taccagtcac gaccatgatt    720
ccagatgttg actgcctgct ctgggccatt gggcggtgcc cgaataccaa ggacctgagt    780
ttaaacaaac tggtaagctg gcttggctctg ccgaaacat ttgtgaatct actgggagtc    840
ttatggtttt attttcccc cagacaccca aaacttgggt ggattccatt ggattctttt    900
ctctttttcc catgtattca gtttgtgatc aaatttcatt gcttcttccct ttgaaattaa    960
aaaaaaaaag tttctnnnnn nnnnnnnnnn nnnnnnnnnn n              1001

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&lt;210&gt; 142

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-436-376 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-436-376.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 10-436-376.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 126..143

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 526..545

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-436-376 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 206,296,317,643..685,853

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 142

```

tccagaagta cactttaaaa tactttgttt ttaaagtgag gcattacaat ggtgtcatga    60
gctgacattc ccagccactg actggaagat ttgggtcaac ccaggggaaca ggcttggttt    120
aaggaggatt aagtgttggg ggcctttggg taaaagttaa tgtctatggt tgtatgtggg    180
attcagtagg gattttcttt aatgtntaaa aattctagtt tcctctgtat ttgctttcct    240
tcttcccttg tcctccctga gcctccacac atgtgtaaca cattttattt gccaantagg    300
tcaaggaggt taaaaangac tttgtcgggc ttggaagtca gcatggttac tgcagttccc    360
ggtaggctac cagtcatgac catgattcca gatgttgact gcctgctctg ggccattggg    420
cgggtcccga ataccaagga cctgagttaa aacaaactgg taagctggct tggctctgcc    480
gaaacatttg tgaatctact rggagtctta tgggttttatt tcccccccag acacccaaaa    540
cttgggtgga ttccattgga ttcttttctc tttttcccat gtattcagtt tgtgatcaaa    600
tttcattgct tcttcctttg aaattaaaaa aaaaagtttt ctnnnnnnnn nnnnnnnnnn    660
nnnnnnnnnn nnnnnnnnnn nnnntcttct ttgtttttga gaaaggtctt gctctgtcac    720
ccaggctgga gttcagtggt gtgatcatag ctcaccgcag tcttaacctc ctagggtcaa    780

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117

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gtgatccacc cacctcagct tctggagtag atggaactat gtgtgtgcca caatgcccag      840
ctaatttttt tantttttatt ttttttagag atgaggtttc accatgttgc ccaggctggt      900
ctcctcggct caagcgattc gcccatctca gcctcccaga gtgctgggac tataggcatg      960
tcccagcatg tcccagaaaa aggccgccac gtctggcctt t                               1001

```

&lt;210&gt; 143

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-431-51 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 10-431-51.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-431-51.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 451..468

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 853..872

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-431-51 potential probe

&lt;400&gt; 143

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agcatccctt atctgaaatg cttgggacca gaagtgtttt cagtttcaga ttttgaata      60
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&lt;210&gt; 144

&lt;211&gt; 980

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

<220>  
 <221> allele  
 <222> 477  
 <223> 10-432-93 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 457..476  
 <223> 10-432-93.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 481..499  
 <223> 10-432-93.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 388..407  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 758..775  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 465..489  
 <223> 10-432-93 potential probe

<220>  
 <221> misc\_feature  
 <222> 297,477,629  
 <223> n=a, g, c or t

<400> 144  
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 aggcaggcat atcacctgag 980

<210> 145  
 <211> 933  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 433

119

&lt;223&gt; 12-631-208 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 414..432

&lt;223&gt; 12-631-208.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 434..453

&lt;223&gt; 12-631-208.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 226..245

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 705..725

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 421..445

&lt;223&gt; 12-631-208 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 30

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 145

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&lt;210&gt; 146

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-260-282 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

120

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<223> 10-260-282.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-260-282.mis2, potential complement

<220>
<221> primer_bind
<222> 220..238
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 636..655
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-260-282 potential probe

<400> 146
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actctcccc tccactctc ccaaggtaac cactaatcta cagttggtgt gtcctcagta      240
aatataggcc agactttcca tgggattcca ttgacaggaa gacaacccgt tcacagggtgc      300
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tcttccccct actggtcccc agtgccttgc tggagcaagc ctatgctgtg cagatggact      420
tcaacctgct agtggatgct gtcagccaga acgctgcctt cctggagcaa actctttcca      480
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cagcgacact ttcttgtcac ctcaggcctc tcacttttgg atgggatggg gtacagactg      960
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<210> 147
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 10-263-26 : polymorphic base A or C

<220>
<221> misc_binding
<222> 483..502
<223> 10-263-26.mis1, potential

<220>
<221> misc_binding
<222> 504..523
<223> 10-263-26.mis2, potential complement

<220>
<221> primer_bind

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121

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<222> 478..496
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 820..837
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 10-263-26 potential probe

<400> 147
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cagtttccac ttgtaatgag atccttggtg tgtcagggag aaaaaggacc tcatagctca      180
tctagtccctg tccctccatt gtacaggcag agggaaacaat atcttgagag ccccagagag      240
aggaatgcag ggacttctgt ctgggggctg ggccctggtag catccatttc tagccagcag      300
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tgattctcag agtcatgttg ctgtatatat gaggtcatgg gcagaggggt cttccaggtc      600
catccaatta ctgaacagcc atctctcttc caacagacat gttctcagtg tcctgagtaa      660
gaccaaagaa gctggcaaga tcctctctaa taatcccagc aagggaactgg ccctgggaat      720
tgccaaagcc tgggagctct acggctcacc caagtaaggg tgtgaaaagg tagcaggagg      780
atcctgcttt agtttcagca ttcattgggtt tagcaacttc tttcttgcc agccatcatt      840
agagaataag gggatttttc taggaataga aacttatacc tttacatgcc aaaattatct      900
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<210> 148
<211> 981
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-258-408 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 10-258-408.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-258-408.mis2, potential complement

<220>
<221> primer_bind
<222> 95..112
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 504..521
<223> downstream amplification primer, complement

<220>

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122

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-258-408 potential probe

&lt;400&gt; 148

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cttccccag	ctttccatct	gaggaccaga	aaagtgtgt	ctcccttaga	tgagatctag	120
acgccccaa	atccttgaga	tgtgggtata	gctcagggta	agctgctctg	aggtaaagg	180
ccatgaaccc	tgccccactc	ctgtcagccc	ctcatcagcc	ttttcagcag	gttccagtgc	240
ctgacttggg	ataggactga	gtggtaggag	gagggggagt	ggaggggcat	agcctttccc	300
taattctgcc	ttaaataaaa	ctgcattgct	gattcagtga	tgattcctta	cttcgtgcat	360
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gtgactggtc	tgtaatcagg	gtccccctag	accagtctct	acaggtggaa	ccctgaagtt	480
tcaatcctta	gccaccact	ratgctctta	ctggatcaca	gggaggaatg	agagtccctg	540
gcaggagccc	aggaggggag	gcaaccaaga	tgggacatac	ataacagttg	tgaactggct	600
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cgctctgccc	aaaccaaaag	ttctagaagg	aagatatttg	ggatagtcct	aggaaatacc	720
cctcccttcc	catctgccac	acaaatcaga	gccactaatg	aataacagc	ctcagggcac	780
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&lt;210&gt; 149

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-317-259 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-317-259.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-317-259.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 742..759

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 297..317

&lt;223&gt; downstream amplification primer.

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-317-259 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 426,432,443,459,841,849..850,898,914

&lt;223&gt; n=a, g, c or t

123

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<400> 149
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gaagtgactg gtagtgagta caaggctgct tttggggggt gatgaagata ttctaaaatt      180
gattgtggta atggctgcac aactctatga atgtcctaaa aatcacgata ccatgggttaa      240
ctaaatgggt gaattgcatg gtatgtgaat tgtacctcaa agctgttaaa aaaaaaaaaa      300
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cagcagtgca cgacagaggc aggattctgc gtgatagttc ttcaggttca cttcacagat      780
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nactatagnn ttttttgccc aagggtacagg agaataaata tgagtagata gaatctcntt      900
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<210> 150
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-323-385 : polymorphic base T or C

<220>
<221> misc_binding
<222> 502..520
<223> 12-323-385.mis1, complement

<220>
<221> misc_binding
<222> 481..500
<223> 12-323-385.mis2, potential

<220>
<221> primer_bind
<222> 868..886
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 416..435
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-323-385 potential probe

<220>
<221> misc_feature
<222> 303,710
<223> n=a, g, c or t

<400> 150
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cgagggtggg agatcacttg aggtcaggag accagcctgg ccaacagggt gaaaccctgt      120
ctctactaaa aatataaaaa ttagccagc atggtggtgc atgcctgtag tcccagctac      180

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124

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tcnttaaaat	ttaagacaaa	acaaaaaacc	ataagaagtg	ggctgggggt	ggtgggacct	360
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gcagttgttc	cccaatgccc	yctgcacagc	aaattcctgt	cctccattac	tagaaataga	540
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aaggcattcc	ataaaattat	cattcttaat	aaaagctagc	aatagaagga	aaacttataa	960
actatatata	gggtatttat	taaaatttaa	cttataggat	a		1001

&lt;210&gt; 151

&lt;211&gt; 857

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 357

&lt;223&gt; 12-324-219 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 337..356

&lt;223&gt; 12-324-219.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 358..377

&lt;223&gt; 12-324-219.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 139..157

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 579..599

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 345..369

&lt;223&gt; 12-324-219 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 442

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 151

attaaagtgt	tgaacaaggc	agcaaagcca	attgccttgg	aagccgccac	cttgaggccg	60
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ccaagctcac	agcaggcagt	gggctgctgg	ccaggtatgg	ccatggcccc	gggggagagt	360
cactacaagg	ggcatcagcg	acctctacca	gccccactgc	ttcagatagg	aagacagagg	420



125

ctcagataag	ctgaggggacc	tnccctcac	cacccaggta	ctaagaggca	ctccccggaa	480
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aatttttact	tttaatttta	cttcagctag	cctgagctgg	gtttctgtca	cacacactcg	600
gtgagcctaa	cacaccaggc	ccagtccctc	cctacagcgg	ctccccaccg	ggcaccacc	660
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ccaatgaatg	tttgccaaag	actgcccaga	ctcccccgct	gtctctaacg	taaaacctgt	840
gcctaaagcc	tggcaga					857

&lt;210&gt; 152

&lt;211&gt; 973

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 473

&lt;223&gt; 12-324-335 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 454..472

&lt;223&gt; 12-324-335.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 474..493

&lt;223&gt; 12-324-335.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 139..157

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 579..599

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 461..485

&lt;223&gt; 12-324-335 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 442

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 152

attaaagtgt	tgaacaaggc	agcaaagcca	attgccctgg	aagccgccac	cttgaggccg	60
gccacgtggg	catctgggtg	agctccctca	gtcattcttg	tctccctgct	ggagacaggg	120
tgtctgatgc	cagcattctt	accctgcatg	acttgccctg	acagcctgcc	tttcatgtac	180
ctttcataatc	cacctgggtt	tcaaatacgg	ccagggacag	agtgcacacg	cgcaccagca	240
tgctggagtc	caccaggagg	ctgctgatct	gcttcaggaa	tacctggaac	tatacagaaa	300
ccaagctcac	agcaggcagt	gggctgctgg	ccaggtatgg	ccatggcccc	gggggatagt	360
cactacaagg	ggcatcagcg	acctctacca	gccccactgc	ttcagatagg	aagacagagg	420
ctcagataag	ctgaggggacc	tnccctcac	cacccaggta	ctaagaggca	ctccccggaa	480
ttcagcacag	atccgacact	ctctccagt	gttttacgct	caaggggtgct	ggattccttt	540
aatttttact	tttaatttta	cttcagctag	cctgagctgg	gtttctgtca	cacacactcg	600
gtgagcctaa	cacaccaggc	ccagtccctc	cctacagcgg	ctccccaccg	ggcaccacc	660
atgctgcgtc	acggcaggcg	cacaagccac	ccattcccac	cctcactccc	taccacaag	720
cagccccggg	tttcatccct	gcattcccag	ggctctagcac	aaagccagac	agagcagggt	780

126

ccaatgaatg tttgccaaag actgcccaga ctcccccgtc gtctctaacg taaaacctgt	840
gcctaaagcc tggcagacca ggactccagg tgaacttctg gacaaccag cacactccac	900
agagccctgg gttaacttta cctttgcatc ggtgttgata tatttattga atctcttgaa	960
tgcatacaacg ttg	973

<210> 153  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-324-380 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-324-380.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-324-380.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 122..140  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 562..582  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-324-380 potential probe

<220>  
 <221> misc\_feature  
 <222> 425  
 <223> n=a, g, c or t

<400> 153	
ggcagcaaag ccaattgccc tggaagccgc caccttgagg ccggccacgt gggcatctgg	60
tgcagctccc tcagtcattc ttgtctccct gctggagaca ggggtgtctga tgccagcatt	120
cttaccctgc atgacttgcc ttgacagcct gcctttcatg tacctttcat atccacctgg	180
ttttcaaate ggtccaggga cagagtgaaca cagcgacca gcatgttgga gtccaccagg	240
gagctgttga tctgcttcag gaataacctgg aactatacag aaaccaagct cacagcaggc	300
agtgggctgc tggccaggta tggccatggc ccggggggat agtcactaca aggggcatca	360
gcgacctcta ccagccccac tgcttcagat aggaagacag aggtcagat aagctgaggg	420
acctncccc caccacccag gtactaagag gcactcccc gaattcagca cagatccgac	480
actctctcca gtggttttac rctcaagggt gctggattcc tttaattttt actttttaatt	540
ttacttcagc tagcctgagc tgggtttctg tcacacacac tcggtgagcc taacacacca	600
ggcccagtc ctccctacag cggctcccac cgtggcacc accatgctgc gtcacggcag	660
gcgcacaagc caccattcc caccctcaact ccctacccac aagcagcccc ggttttcatc	720
cctgcattcc cagggtctag cacaaagcca gacagagcag gttccaatga atgtttgcc	780
aagactgcc agactcccc gtcgtctcta acgtaaaacc tgtgcctaaa gcctggcaga	840
ccaggactcc aggtgaactt ctggacaacc cagcacactc cacagagccc tgggttaact	900
ttacctttgc atcgggtgtg atatatttat tgaatctctt gaatgcacac acgttgaact	960
tcatcagctc cccagggagg tcaaagtaac tctggagcac a	1001

<210> 154  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-325-30 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-325-30.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-325-30.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 472..491  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 926..945  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-325-30 potential probe

<400> 154  
 ctttgggagg tggaggcgtg tggatcacao ggtcagaaga tcaagatcat cctgggctaac 60  
 acggtgaaac cccgtctcta ctaaaataca aaaaattagc caggcgtggt ggtgtgtgcc 120  
 tgtaatccca gctactcagg gaggctgagg cgggagaatt acttgaaccc aggaggcgga 180  
 ggttgacgtg agccaagatc gagccactga actccagcct aggcgactga gtgagactcc 240  
 atatcaaaaa aaaaaaata caaagcctca acccctcctt cccatcaggc ctcttgcac 300  
 agagtctctg ggatggggcc caggaatctg tattctttcc cagctcccca gaatgttcag 360  
 ccaggtttgg aaactgatct atccgattct tcttggttca cagttaggga atctgtagct 420  
 ctgggaaggg aaggaacttg cccagtcac atctgatatt agtgcttctt tctccaatga 480  
 agagccttta ggctgggagt ycagagacat gggttcaagt ccaggctata ccagtcacatca 540  
 cctcgggcaa gtcatttcac ctctccaage ctctgcttcc ttactgtgag aataatgcca 600  
 ttgtgttggg aatcaaaaaga gagagtggca atggaaatgc tttgtcaagc tttctatattt 660  
 gtgcacatgg aagttgttaa gagctagaac cagccagtgt tcaactcctgt ataccacgct 720  
 gttcccttcc aacagaggtc agggctctgc tgtgttgggg gtggccgcca gccagtttcg 780  
 gtggttgctg ggcttcaggc catctgttac caactctctt ctctccatct tttgcagggtg 840  
 ttgggatggc caccaactgg gggagcctct tgcaggataa acagcagcta gaggagctgg 900  
 cacggcaggc cgtggaccgg gccctggctg agggagtatt gctgaggacc tcacaggagc 960  
 ccacttcctc ggaggtaagc ccctagctcc tccccacagc a 1001

<210> 155  
 <211> 999  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 479

128

&lt;223&gt; 12-327-31 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 459..478

&lt;223&gt; 12-327-31.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 480..498

&lt;223&gt; 12-327-31.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 449..468

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 982..999

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 467..491

&lt;223&gt; 12-327-31 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 758

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 155

tgtcttataaa	gctagttttt	caatgttaac	aggctggaaa	tattgttaat	acaaaaacct	60
atcactgtgt	accttatggg	acaccaaata	aacaaggtaa	aattatacaa	atgtatcatt	120
aaaaagcagc	agtagaaaac	catgaccatg	cagaggtagc	cctaatacatg	ctctgaaaac	180
agttctttgt	ttctaaagtc	cagttgtgtg	cattccccag	gctggctctg	cagagttatc	240
aagtgtctta	gagcagcctc	ctctcctgag	gctgaatatg	aacctgccat	tcactctgtt	300
attgtctagt	tttagttagg	aacatgaggt	gatcataact	atactttgta	ggttattagg	360
gaatataatt	ttacatgttg	tagtcatatg	taaaggtaga	agtttgtgga	ctccaacccc	420
agtttattct	ctctctagat	ttgcttttcc	tcctgtgtca	ctagatacaa	ggctcttckt	480
ggctctgtga	cggtttttct	cccactcctt	tatgatattg	ggggagttct	catttttagga	540
aatttacatt	tttaaaaaat	atgtgacttt	ccaacatggc	acatatatac	atatgtaaca	600
aacctgcacg	ttgtgcacat	gtaccctaga	acttaaagta	taattttaaa	aaaagtgact	660
ttcattataa	ctaaattata	cctcagagct	ggtgtacaca	cctccatata	cattgacaaa	720
aggtagtatt	tgctaagctc	ctggatatgtg	gcaaacgnct	gggtaactgc	tttacctaga	780
agtgatttta	tttaatcctt	caagtgcctt	gtggaataga	gattagtatc	cccattgcac	840
atgtgaggaa	actgagggct	agaaagggtc	tgtattggat	gcctgtgtat	gccgaggcac	900
taagtaactt	tcattcttta	gtttcccat	taaggaatct	attctggcag	atgattttga	960
tcctgttagt	attgattgct	tgcttcagat	gcctatttg			999

&lt;210&gt; 156

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 500

&lt;223&gt; 12-327-415 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

129

<222> 480..499  
 <223> 12-327-415.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 501..520  
 <223> 12-327-415.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 86..105  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 615..633  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 488..512  
 <223> 12-327-415 potential probe

<220>  
 <221> misc\_feature  
 <222> 395,958,975  
 <223> n=a, g, c or t

<400> 156  
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 ttattctctc tctagatttg cttttcctcc tgtgtcacta gatacaaggt cttccttggt 120  
 ctgtgtacgg ttttcttccc actcctttat gattttgggg gagttctcat tttaggaaat 180  
 ttacattttt aaaaaatatg tgactttcca acatggcaca tatatacata tgtaacaaac 240  
 ctgcacgttg tgcacatgta ccctagaact taaagtataa ttttaaaaaa agtgactttc 300  
 attataacta aattatacct cagagctggt gtacacacct ccatatacat tgacaaaagg 360  
 tagtatttgc taagctcctg gtatgtggca aacgnctggg taactgcttt acctagaagt 420  
 gattttatatt aatccttcaa gtgccctgtg gaatagagat tagtatcccc attgcacatg 480  
 tgaggaaaact gagggctagr aagggtctgt attggatgcc tgtgtatgcc gaggcactaa 540  
 gtaactttca tcttttagtt tccatttaa ggaatctatt ctggcagatg attttgatcc 600  
 tgttagtatt gattgctttc agatgcctat ttgtaaaactg acttaagtaa aaccagcatt 660  
 acccattatt cttgaggaat gaactgttct ggtcagctgg gcttttttga ttaactgaga 720  
 acggaaagcc cagtttttgt ttttgttttt ttatttgga gttttttctt aaagtctcta 780  
 taaaataatc tagattcact ttcatactcg tgtgactagc ctagtagaca gtctatgtgt 840  
 gattctgctt gtagcttttg ggataaagcc taaaagttgt agttccacac atgaggttgt 900  
 gcaggatctg acccttccca accctgacca taattcctac tccattccct taccgcgnac 960  
 gattttctcc attcnaaatg ccctgggaag gccaaagctgg 1000

<210> 157  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-331-270 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-331-270.mis1, potential

<220>

130

<221> misc\_binding  
 <222> 502..521  
 <223> 12-331-270.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 751..770  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 285..305  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-331-270 potential probe

<400> 157  
 aactacggct tgggaggtgt actaagccaa gggccagaac acaaaatcag gtcctaacac 60  
 taagtgccat cctcttttgt ttctaccaga ttgcttttctt tattcaccat tcttcctggc 120  
 tcagtgggtca acctagtaga gtcttgctca atatatgggc ggggcttctg gttctcattt 180  
 tgtttttctaa caaggaaaat gaagaaatag gcataatgga gttaaaatga gtcttaaagg 240  
 tcacagtctg ccaacacttg tcaccttttg acaagtgtcc agtttctact tggacagctc 300  
 taaggccaaa ggagtttact acctgctgca gcagtcctgg ctcttaacaa ctgctacaaa 360  
 attctgcctc cccgtaattt ctaccagtta gtctgtattt tcaggcacac ataaaataag 420  
 cccgattccc ccaactgtctg aaagcagctc ttatgtctgc cattctactc gaattattccc 480  
 atcttaaaca ttgcagacct rgtaggggag ataaacagag cccagtggc aggggtactt 540  
 acacagacac aggtattata caggatactc aagcagtcct tggacatttc atcacttact 600  
 gaaagtgaag gggtaatgac tgggtatttt tctcaatgaa aagatagtag aacagttacc 660  
 acattttttt cagcataaga actgttataa tccccccaa gatgattatt ttccctttaa 720  
 tatacttctt ccagcccctg tctaccaaag gttttcatag tgcccacag aacctacaag 780  
 ccattttgca tattgtcttt gggttgatag agcagctgtt ttcttattat gctccacagt 840  
 cttcataatt tcaacctgtt gttgtgccat catttaataa agccattttc ccttaatgtc 900  
 ttgcttagga tatatctagg atttggtttt tttgtttttt gtgttttttg gctatttaag 960  
 tagaccacaa atgaacaact taatatatat gtgtatcctt t 1001

<210> 158  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-331-275 : polymorphic base T or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-331-275.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-331-275.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 756..775  
 <223> upstream amplification primer, complement

131

<220>  
 <221> primer\_bind  
 <222> 290..310  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-331-275 potential probe

<400> 158  
 taagaaacta cggcttgga ggtgtactaa gccaagggcc agaacacaaa atcagggtcct 60  
 aacactaagt gccatcctct ttgttttcta ccagattgct ttctttattc accatttcttc 120  
 ctggctcagt ggtcaacctt gtagagtccct gctcaatata tgggcggggc ttctgggtctt 180  
 catTTTgttt tctaacaagg aaaatgaaga aataggcata atggagttaa aatgagtctt 240  
 aaaggtcaca gtctgccaac acttgtcacc ttttgacaag tgtccagttt ctacttggac 300  
 agctctaagg ccaaaggagt ttactacctg ctgcagcagt cctggctctt aacaactgct 360  
 acaaaattct gcctccccgt aatttctacc agtttagtcct gatttttcagg cacacataaa 420  
 ataagcccgga ttccccctact gtctgaaagc agctcttatg tctgccattc tactcgaata 480  
 ttcccatctt aaacattgca kacctggtag gggagataaa cagagcccca gtggcagggg 540  
 tacttacaca gacacaggta ttatacagga tactcaagca gtccttggac atttcatcac 600  
 ttactgaaag tgaaagggtta atgactgggt atttttctca atgaaaagat agtacaacag 660  
 ttaccacatt tttttcagca taagaactgt tataatcccc tccaagatga ttatttttcc 720  
 tttaatatata ttcttccagc ccctgtctac caaagggttt catagtgcc atcagaacct 780  
 acaagcccat ttgcatattg cttttgggtt gatacagcag ctgttttctt attatgctcc 840  
 acagtcttca taatttcaac ctggtgttgt gccatcattt aataaagcca ttttccctta 900  
 atgtcttgct taggatatat ctaggatttg ttttttttgt tttttgtgtt ttttggctat 960  
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<210> 159  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 12-334-320 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 12-334-320.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..523  
 <223> 12-334-320.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 184..202  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 616..634  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 12-334-320 potential probe

```

<400> 159
ccagcctggg caacagagtg agactccatc tcaaaaaaaaa gaaaaggata acgtgataga      60
cttatagggt ggggcagcct ccagggatga aacatctgaa tgaccaaaagg agccagtcac      120
gccaggattt tggaggaaaag caccagggca gagagtgcag aaagggcaaa cgctccctgg      180
aaagattctt agtcaagagt ccttcactcc cagtcctacc acaaactggg tcaccttgaa      240
caagtcacgt aacttctgag gctcagctgc cacatctaca aaatgggaat aaagacatct      300
tacctgccac attgtgagag gtttcaacca aagggctggt aaggtctggg atcctcccca      360
aatctcacca tagacacctg atactcatca cttggcaccc gtcttggaag aggggaacct      420
gcacagagaa ccctgggtca tgcttttgat ttttaatttc atgctgcact agaaatagct      480
tcttttgttc ctggttgacc cargagcctc tcctgccac ctggggccta ttctagttaa      540
cagctgctta tcccctcagg tacaaaagcc aacaaggaaa ggacatcagg aaacattggt      600
ctgggaataa ccagacacct atctgccacc atctccccc atcccgtgac cacacacggg      660
agactggagg actcagcctg tcctgtagtc agataatgta catggtttat ttaaagagtc      720
aaaaggggcc gggcgcagtg gctaacgcct gtaatcctag cactctggga ggctggggtg      780
ggtggatcac ctgagctcaa gagtttcaga ccaatctggc caacatggtg aaaccctgtc      840
tctacaaaaa atacaaaaat tagccgggtg tggtggtgga cgctgtaat cccagctact      900
tgaggaggctg aggcaggaga attgcttgaa cctgggaagt ggaggttgca gtgagctgag      960
atcgtgccac tgcactccag cctgggcaac aacaacgaaa a                                     1001

```

```

<210> 160
<211> 1001
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele
<222> 503
<223> 12-334-391 : polymorphic base A or G

```

```

<220>
<221> misc_binding
<222> 483..502
<223> 12-334-391.mis1, potential

```

```

<220>
<221> misc_binding
<222> 504..523
<223> 12-334-391.mis2, potential complement

```

```

<220>
<221> primer_bind
<222> 113..131
<223> upstream amplification primer

```

```

<220>
<221> primer_bind
<222> 545..563
<223> downstream amplification primer, complement

```

```

<220>
<221> misc_binding
<222> 491..515
<223> 12-334-391 potential probe

```

```

<400> 160
gggcagcctc cagggatgaa acatctgaat gaccaaagga gccagtcacg ccaggatttt      60
ggaggaaaagc acccaggcag agagtgcaga aagggcaaac gctccctgga aagattctta      120
gtcaagagtc cttcactccc agtcctacca caaactgggt caccttgaaac aagtcacgta      180
acttctgagg ctcagctgcc acatctacaa aatgggaata aagacatctt acctgccaca      240
ttgtgagagg tttcaaccaa agggctggtt aggtctggga tcctcccca atctcaccat      300
agacacctga tactcatcac ttggcacccg tcttggaaga ggggaacctg cacagagaac      360
cctgggtcat gcttttgatt ttttaatttc tgctgcacta gaaatagctt cttttgttcc      420

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133

tggttgaccc	aagagcctct	tcctgccacc	tggtggcctat	tctagttaac	agctgcttat	480
cccctcaggt	acaaaagcca	acraggaaag	gacatcagga	aacattgttc	tggaataaac	540
cagacaccta	tctgccacca	tctcccccca	tcccgtagcc	acacacggga	gactggagga	600
ctcagcctgt	cctgtagtca	gataatgtac	atggtttatt	taaagagtca	aaaggggccc	660
ggcgagctgg	ctaacgcctg	taatcctagc	actctgggag	gctgggggtg	gtggatcacc	720
tgagctcaag	agtttcagac	caatctggcc	aacatgggtg	aacctgtct	ctacaaaaaa	780
tacaaaaatt	agccgggtgt	ggtggtggac	gcctgtaatc	ccagctactt	gggaggtgga	840
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gcactccagc	ctgggcaaca	acaacgaaaa	ctccgtctca	aaaaaaaaaa	aaaaaaaaaa	960
aagagtcaaa	aggatcttgg	tccctgggtt	gggccactga	t		1001

&lt;210&gt; 161

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-335-417 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-335-417.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-335-417.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 85..102

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 534..552

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-335-417 potential probe

&lt;400&gt; 161

agcctttgct	gcttttgttc	ctgcaatttg	gaacactgtc	cccatcccag	cctctcacct	60
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134

cactgggtcg catcgaggcc ccgccccctg agcctgggta g

1001

&lt;210&gt; 162

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-337-189 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-337-189.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-337-189.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 313..331

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 792..812

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-337-189 potential probe

&lt;400&gt; 162

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gggcaactag tgtctagtga ggggggttggg ctggcgcgca ctgatcccag actttccgga      60
tcttctgcct ttagatcggg ccggtgtcgg ggcattgtagg ccagtgagac tggagccagt      120
tagagctaca acggggagcg attagggcca aactttgtcc aggggtggaag cgagcggggc      180
cgtgaagtgg ggccagcctg ggcagccgac cgtgtcgttg cctcggggcc tttccaggca      240
ctggcctaag tcctggcgat aaagtgcgac cgatttcctt gtgggcggtt tgaggctttc      300
ggtgatctga cccgtctgtc attcattctt cattcattca tgtgatgaat gaatacagta      360
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gagactcgca ggaaaaaatt ggatattcta gtttgaagg aggaaagtat tgctgtgaag      900
atgtagattt gaatgtcatc agcaaaacat aaataaagcc aaggggagggt tgaggctgta      960
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&lt;210&gt; 163

&lt;211&gt; 881

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

135

<222> 381  
 <223> 12-340-130 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 361..380  
 <223> 12-340-130.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 382..401  
 <223> 12-340-130.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 252..272  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 681..701  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 369..393  
 <223> 12-340-130 potential probe

<220>  
 <221> misc\_feature  
 <222> 205,247,340,499,507  
 <223> n=a, g, c or t

<400> 163  
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 ctccagcctg tgacacagtg caactctatc tctaaaatca tcccaacaat ttttaaaagc 180  
 tgtaattaaa aaaaattctg ttaanacaaa ttctagggtta atataaacag aaaaaataa 240  
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 agacagtttt tcacattttc cttgacaaac accttttccn ttttaaaatg agaaaaatta 360  
 ggacctttgt tttccaaaaa rgtccaattt tcacttaaag catttaaaaa ttatctatat 420  
 gctgagaata tgactaagcc catatgttta aagacattcc ctatatacgt atgtattttt 480  
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 agccccctct agctataggc caatacttag gcagcatgtc caagacacct caagccaaat 600  
 gaagaaggaa gattccatag cattttttta attaaaaagc cagcaggcaa atctttgaga 660  
 ctagacagtt cagcttgagg gtctgagaaa gcctctgctc atctcaaacc agcaacaaat 720  
 ctttagaaaag taattccat gccatgagat tgctcgtggc atgaatgtga cactataatt 780  
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 aatcctaaac tactacacca cagattgtca attctagaga a 881

<210> 164  
 <211> 961  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 461  
 <223> 12-340-210 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 442..460

136

&lt;223&gt; 12-340-210.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 462..481

&lt;223&gt; 12-340-210.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 252..272

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 681..701

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 449..473

&lt;223&gt; 12-340-210 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 205,247,340,499,507

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 164

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gctgagaata	tgactaagcc	catatgttta	aagacattcc	ytatatacgt	atgtattttt	480
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caacatcctg	aattaagaga	gatgtgttat	tttagcttaa	agcagcagat	taaaaaataa	840
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gcacatctgt	atttattgtt	aactgcactt	gcaggaggga	aaagaaagaa	gtctcatgac	960
t						961

&lt;210&gt; 165

&lt;211&gt; 973

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 473

&lt;223&gt; 12-340-222 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 453..472

&lt;223&gt; 12-340-222.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

137

<222> 474..493  
 <223> 12-340-222.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 252..272  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 681..701  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 461..485  
 <223> 12-340-222 potential probe

<220>  
 <221> misc\_feature  
 <222> 205,247,340,499,507  
 <223> n=a, g, c or t

<400> 165  
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 ggacctttgt ttcccaaaaa agtccaatth tcaacttaaag catttaaaaa ttatctatat 420  
 gctgagaata tgactaagcc catatgttta aagacattcc ctatatacgt atktattttt 480  
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 gaagaaggaa gattccatag cattttttta attaaaaagc cagcaggcaa atctttgaga 660  
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 ctttagaaaag taattcacat gccatgagat tgctcgtggc atgaatgtga cactataatt 780  
 caacatcctg aattaagaga gatgtgttat tttagcttaa agcagcagat taaaaataaa 840  
 aatccctaaac tactacacca cagattgtca attctagaga agcactgggc tcagctttct 900  
 gcacatctgt atttattgtt aactgcactt gcaggaggga aaagaaagaa gtctcatgac 960  
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<210> 166  
 <211> 975  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 491  
 <223> 12-340-240 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 471..490  
 <223> 12-340-240.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 492..511  
 <223> 12-340-240.mis2, potential complement

<220>

<221> primer\_bind  
 <222> 252..272  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 681..701  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 479..503  
 <223> 12-340-240 potential probe  
  
 <220>  
 <221> misc\_feature  
 <222> 205,247,340,499,507  
 <223> n=a, g, c or t

<400> 166  
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 ctccagcctg tgacacagtg caactctatc tctaaaatca tcccaacaat ttttaaaagc 180  
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 aatcctaaac tactacacca cagattgtca attctagaga agcactgggc tcagctttct 900  
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<210> 167  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens  
  
 <220>  
 <221> allele  
 <222> 501  
 <223> 12-341-99 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-341-99.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-341-99.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 403..422  
 <223> upstream amplification primer

139

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<220>
<221> primer_bind
<222> 927..947
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-341-99 potential probe

<220>
<221> misc_feature
<222> 133,141,738,898
<223> n=a, g, c or t

<400> 167
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aatatgacta agcccatatg tttaaagaca ttccctatat acgtatgtat tttttattat      120
tgtttaaadc aangctagga ngctacccca gaacaaaaag aaaaattctt ccttagcccc      180
ttctagctat aggccaatat ttaggcagca tgtccaagac acctcaagcc aaatgaagaa      240
ggaagattcc atagcatttt ttaaattaaa aagccagcag gcaaattctt gagactagac      300
agttcagctt gagggctctga gaaagcctct gctcatctca aaccagcaac aaatctttag      360
aaagtaattc acatgccatg agattgctcg tggcatgaat gtgacactat aattcaacat      420
cctgaattaa gagagatgtg ttatttttagc ttaaagcagc agattaaaaa taaaaatcct      480
aaactactac accacagatt rtcaattcta gagaagcact gggctcagct ttctgcacat      540
ctgtatttat tgtaaactgc acttgcagga gggaaaagaa agaagtctca tgacttgagg      600
caacaatgaa aactgccctg aacatatgcc tgctttgctt tgtataatag agacctaagg      660
tcacatccta gaaaagggtga agtaattatt atacagtata taaccatttt atagcctggt      720
ttcatcatga attttcnca tattactata actttaacat tatattaaca acttcataaa      780
attcaacaaa atagatagaa taaccattat cttctcctgg tagatagtca ttacaaataa      840
tgttacacct aacgggtcttg gtatacaca gctttatctc attttgactt acgtcttnca      900
tcagagtcac agaaatggaa acaacagggc aaaaatgagta gtaatgttta aagctctgaa      960
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<210> 168
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-342-32 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-342-32.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-342-32.mis2, potential complement

<220>
<221> primer_bind
<222> 513..532
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 81..101
<223> downstream amplification primer

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<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-342-32 potential probe

<400> 168  
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 aaaaactatg gaaaaaatatc actttttctca taactgctta gtaaataactt aaaatacact 120  
 gtttgtggaa gggggcttct ggggtactggg aatgttctat ttcctaattt ggggtgtagt 180  
 gcaaaagtga atgtattttg tggttaattat tgggtgtactt ttctgtatga aggttatact 240  
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 aaacagagcg tcatactttg ttcctgggga tataaactga caaaacttct taggttaggta 360  
 aacttcagat ttataaacia taaaaaacat acgttcttgt tccaaaggta tatactcaca 420  
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 gaaaaattat aaatgccagt yaactggggc ctgggtgaaat aaattatggc acaatcacac 540  
 caggaaacac taaccattta attttttttt gagagctcat caccagggt agagtgcagt 600  
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 gcctcctgag tagctgggat tacaagtggc tgccaccgtg cccatctaatt tttgggtattt 720  
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 tccacctgcc ttggcctccc aaagtgtctg gattacaggc gtgagccact gcgcccggcc 840  
 taaaattgtt ttaattaaaa aaaaaaaaag aagaagaaa aaggtagaga atagtgtagc 900  
 cgtgtatatg gtatatcact ctatgtgcgg agaaaataaa ctaagggtgga ggtgggtggg 960  
 aatatttaca cacacacaca cacacacaca caaaaaaac a 1001

<210> 169  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-344-349 : polymorphic base G or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-344-349.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-344-349.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 154..173  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 685..705  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-344-349 potential probe

<400> 169  
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 accacaaatg aacaacttaa tatatatgtg tatecttttc atgtttgtat taaactagtt 120



141

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cctcaagata cattttcaaa agtagaacta ctagaatcta aagtataggc atcttttttt 180
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agctcactgt agtctcgac tcctaggctc aagcaatttt cctcctcac cctcccaggt 300
ggctaggact gcatgcgcgt gtaccacgcc tggctaaaag ttaaaagtat aagcatctca 360
attgctcttg ttataaatcc ccattgaaag ttctactgca gcaaacaaaa ataccatttt 420
tatcagactc attaaactgag catttctttt tttgttgctg ttatctgtgt tgggcaaatg 480
gctcttaaatg tctgatctat kactaattct tataaagaaa ctactacttc cctgtgagca 540
ggaacaggaa agtagcataa aaacgaaact taattttcat gagcccaaaa ccaaaagatg 600
atgaaactga gtatgaacca tggacaaagt gctttatttg gcacttgctt ctatttttta 660
aaattgtttc cattcactcc taagctgcta ggaatgctaa aatgctaggt ctcttctca 720
gatcccaatg ggcagccctg ccaagctagc ttgactagat ggggtgtctc ataactctga 780
gacaacgcag acgctctgca tcccttatca ccacctttca acctactggg gctctgtgat 840
tgatttattt aacgacttca gagcagggtg aaacactgct ctgcatgtct cactatatat 900
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&lt;210&gt; 170

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-345-453 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-345-453.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-345-453.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 53..70

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 558..578

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-345-453 potential probe

&lt;400&gt; 170

```

cagatcccaa tgggcagccc tgccaagcta gcttgactag atgggttgct tcataatcct 60
gagacaacgc agacgctctg catcccttat caccaccttt caacctactg gggctctgtg 120
attgatttat ttaacgactt cagagcaggg tgaacactg ctctgcatgt ctactatat 180
atctcactat ttctagatcc tccattttta aaaggaattg ggattcactc attcatcaag 240
tattgttcag catctactct agagcagata cagtgggtgag ggaaaaaaag tctcagcctt 300
catggaacct ataaatctag tgggaagaag agagactaaa caagtgcacc aacaaataat 360
tacaatatgc tacttgctat gagggatcat agagaacatg aaagaccaag ttagaatata 420
cagctgagac attaaaaatg aatataaatc ctcttaataa ccactactgc atgcacaaat 480
gaggccttat acagtatgtc statgtgggt aaaaaaaaaa tcttttaaat taaaaaacg 540
caaagtgggt aggaggcat tcaccaaaca ttaactgaag tcatctctgg cttttaacac 600
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142

tcatgcctgt	aatcccagca	cttgggagggc	tgagggtgggc	agatcacttg	agggtcaggag	720
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aaaatcactt	gagctcagga	ggcggagggt	gcagtgaagg	aagatcatgc	cactgcactc	900
cagcctgggc	aacagagcga	gagactgtct	caaaaaaaaa	aaaattgaat	acctaaaaag	960
aattatttgt	aacgtgtaag	ttataaaaaa	taataataga	a		1001

&lt;210&gt; 171

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-346-204 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-346-204.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-346-204.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 684..704

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 248..267

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-346-204 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 664,706

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 171

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ccccatcaca	gagattacaa	ggggaagtg	ctagagggcc	tcaggcagaa	tttttttttt	120
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tgccccatga	gcaatcgaaa	gagaccttaa	gaaagtattt	taaaaatgat	taaaaacaga	360
atatgcagca	gaaaaaagat	attaacatca	tggctaagca	gtttccgaat	tgagccagac	420
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gccggatgtg	cagtggcttg	atcttagttc	actgcaacct	ctacctcccg	gggtcaagtg	900

143

attctccac ctcagcctcc tgagtagcag ggattacagg tgccccacca acatgcctgg 960  
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<210> 172  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-364-55 : polymorphic base G or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-364-55.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-364-55.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 447..464  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 849..867  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-364-55 potential probe

<220>  
 <221> misc\_feature  
 <222> 180  
 <223> n=a, g, c or t

<400> 172  
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 actgtgtgcc tggtagcttc tggactaaac aagcatgagt ctggaggata attattaatn 180  
 cctgcagaga tgaggaggagg aggagcaagg tattggggct ccagagatgc ctgggcaaag 240  
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 ggagcagggg ttgagcagga gacagaactg cattctcgag caggagccaa agaattgata 420  
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 ttctaagccg gatcataatt acatggatgc catgggattt ggaatgggca attgctgtct 900  
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<210> 173

144

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<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-364-108 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 10-364-108.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-364-108.mis2, potential complement

<220>
<221> primer_bind
<222> 394..411
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 796..814
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-364-108 potential probe

<220>
<221> misc_feature
<222> 127
<223> n=a, g, c or t

<400> 173
atatttttact ctcacagcat gtgcatgttt tcattaggac aagccatatt tcaagtactc      60
agtgccact gtgtgcctgg tagcttctgg actaaacaag catgagtctg gaggataatt      120
attaatncct gcagagatga gggaggagg agcaaggat tggggctcca gagatgcctg      180
ggcaagtgg gtactaaagt ttgaccaaag tctttggagt taatttggga catgttgcaa      240
acagtgaag cagaccatgg cagactttga acacctgaat tgatttggta agttacaggg      300
agccttggga gcaggggttg agcaggagac agaactgcat tctcgagcag gagccaaaga      360
attgatacta atgcaaaatc tttaatcaca aagtccctt gtagtgtctt gacaccacag      420
atacattttc tgttgagttg tcttagagaa ctaatttgag gtatgatttc attccactta      480
cttgtagtga atcaaattct ycttttacac gatgactctt aggtgaccaa tttatgacgt      540
ttggttgtct gtttttagtac cttaacaaga aatatccgac ataggagagg agaaaagggt      600
gtcatcaatg taccaagtaa gtctactgag aggtgggtggg gtgggagaga gacatgttgt      660
attgttggtt aatcctggat tctaaacat ttttattttt gtatttttat aatacagtat      720
ttaaggacaa gaatacacca tctccattta tagaaacatt tactgaggat gatgaagctt      780
caagggcttc taagccggat catatttaca tggatgccat gggatttgga atgggcaatt      840
gctgtctcca ggtatagttt caaatataca gagaggcaaa gtgttccatc catttctggt      900
ttttaacttc ttatatatg catgtttcct gttccaaaaa tcacatttta atgaggttga      960
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<210> 174
<211> 1001
<212> DNA
<213> Homo Sapiens

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145

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<220>
<221> allele
<222> 501
<223> 10-364-267 : polymorphic base A or T

<220>
<221> misc_binding
<222> 481..500
<223> 10-364-267.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-364-267.mis2, potential complement

<220>
<221> primer_bind
<222> 235..252
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 637..655
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-364-267 potential probe

<400> 174
ttggggctcc agagatgcct gggcaaagtg ggtactaaag tttgaccaa gtctttggag      60
ttaatttggg acatgttgca aacagtgaaa gcagaccatg gcagactttg aacacctgaa      120
tgattttggg aagttacagg gagccttggg agcagggggt gagcaggaga cagaactgca      180
ttctcgagca ggagccaaag aattgatact aatgcaaaat ctttaatcac aaagtccct      240
tgtagtgtct tgacaccaca gatacatttt ctgttgagtt gtcttagaga actaatttga      300
ggtatgattt cattccactt acttgatatg aatcaaattc ttcttttaca cgatgactct      360
taggtgacca atttatgacg tttgggtgtc tgtttttagta ccttaacaag aaatatccga      420
cataggagag gagaaaaggt tgtcatcaat gtaccaagta agtctactga gaggtgggtg      480
ggtgggagag agacatgttg wattgttgtt taatcctgga ttctaaacca tttttatttt      540
tgtattttta taatacagta ttaaggaca agaatacacc atctccattt atagaaacat      600
ttactgagga tgatgaagct tcaagggtct ctaagccgga tcatatttac atggatgcca      660
tgggatttgg aatgggcaat tgctgtctcc aggtatagtt tcaaataac agagaggcaa      720
agtgttccat ccatttctgt tttttaactt ctttatatat gcatgtttcc tgttccaaaa      780
atcacatttt aatgaggttg aaatggtagc tggatatgct ttctgaaaaa caatgaagtt      840
atattagtaa attcattgga agctgtctat gactaatagt tctacagact ctgttgttca      900
ccacaaaggt atatacggta tatatacctt tataactgta atttcagtta acttaaaatg      960
caatatattc tgtcattgtt tcccttctct ttttatatgc c                                1001

<210> 175
<211> 997
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 497
<223> 10-367-20 : polymorphic base G or T

<220>
<221> misc_binding
<222> 477..496
<223> 10-367-20.mis1, potential

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<220>  
 <221> misc\_binding  
 <222> 498..517  
 <223> 10-367-20.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 478..495  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 889..908  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 485..509  
 <223> 10-367-20 potential probe

<400> 175  
 agcgttggtta acttgcccct ctctgccag caccttcctt ctctgacttc actgcttggg 60  
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 aagcctgcag tatactctgag gccagatacc tttatgatca gttggctact atctgtccaa 180  
 ttgttgtaag tagaaattac ctcttatttt aaatactact tcgtatgaaa taagatagca 240  
 tgtgcagaat ttactgacag tgtgctattt aagtcagtt aagacctcag tcagagatgg 300  
 actaatataa atagtatgta ggtttaggta taatgaactg agagtctaca ctgtagaagt 360  
 ttactcttgc tagtacaaca ttgatttggt aaatgtgaag tttgaatgtg gccattttcc 420  
 ctccccata cttcatgtcc tcacattaga gagaggatga ttttaagtga tcaaacccaa 480  
 ggaactggat tcttcckggt atatttcact gtaagatagg cacaggatca tgtgctctgt 540  
 atggggtgca taaacatgct ttttgatcag aaatataact gcatggagct ttttttagca 600  
 tgtaagtgtc actttgaatt tgcaggagct gacttttgtt tgtttttaga tggctttgag 660  
 tgctgcatct cccttttacc gaggctatgt gtcagacatt gattgtcgtt ggggagtgat 720  
 ttctgcatct gtagatgata gaactcgga ggagcgagga ctggaggtgg gaattgtttt 780  
 tccttaatat cccttttaag tcaagcaggt aaaatggatc ttttgtaact acttgcaatt 840  
 tcaggatgtc tccctgcaat ttttatctga aatggggaaa aagatttggg ctagggtggg 900  
 agttaatttt tatcgtatgt tgatgtctgt atttgtttta gcttcattta aaattagtag 960  
 tccgattttt ctatattttg gaagtgtctg tggctgt 997

<210> 176  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-351-389 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-351-389.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-351-389.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 113..132

147

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 518..537

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-351-389 potential probe

&lt;400&gt; 176

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gtgatctgct tttaaaagat aaaagtatat ttaaatacata ccagaaaaaa agggatattta      60
aattgccttc agatttttaa aaacataatt ttccttaata atacttttta gggctcacta      120
acttttttct tgtatcttat gttgagggtt ttatataatt atcatatata aaaatatatt      180
ccactcattt tatatagcat attcatttat atatatattt aatcttttagc cattgaagaa      240
caataactat aggatcagta aatcccgata tgactcaata gacagctatt tatctaagtg      300
tggtgagaaa tataatgaca tcgacttgac gatagataaa gagatctacg aacagctgtt      360
gcaggaaggt ggggtttctac tccatcttct tgggtttgaa tgtacgcctt tagttcttca      420
aagctctttt acactttttt gctgactcct ggtctgggtt ctatttttag tgcaaagtct      480
ttaacttctc tcatgaggct rcttgttcct aaagtttcag gttccaaata cttgtgagat      540
tttcttgatt tttagcaaaa ggagcttact ttggagggtc gtgtctaggt ccacaccagt      600
gccccagcag gcattgtgta agtagtgaat aaaacgctga cactcccga gtgtccgcag      660
gataaaatac tgggtgaggt gaggatacta atcaaaccag ctatttcaag cttctagaca      720
ccctttcttc aaacttcctg agctatcctg tcgtctttct ggactaagag gaagtgggtat      780
ctctcgcttt gcagtgaagt cccatgtgat ctctttttgc aggcattgat catctcctgg      840
cccagcatgt tgctcatctc tttattagag acccactgac actgtttgaa gagaaaatac      900
acctggatga tgctaataag tctgaccatt ttgagggtat ttctcagttt tatttttatt      960
tatttatgta ttgacttctg agagtctgga atcctctggt t                                1001

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&lt;210&gt; 177

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-353-102 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-353-102.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-353-102.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 400..417

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 800..819

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

148

&lt;222&gt; 489..513

&lt;223&gt; 10-353-102 potential probe

&lt;400&gt; 177

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ctttccccc ccaaactcac aaaaggagat ataaatecctt ccagtcttgc catcgggtcc      60
caagatgggc aggacggaga aagaactctc tctcagtgtg ttgagaaaca gctacccttc      120
ttaggctcctt gggtcacgga agcagcagca ttcccaggag aggaggaaga aaggaaacaa      180
ctgtcatgtg acagggttct cagatgaaag tgtccagtgt cacctggaaa ggcaggctgg      240
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gctgggagat agggagtagg agctgggtga gattaagctt gcagtgatag ggcctcatag      360
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catgcatcag caagaactga ctcaaaagta ctctttcgcc agctgttcat cacccttctg      720
atgctgctat gagaaggctc ttatatataa aataaaaatt aagtatgtgc atggtagcaa      780
gtcagtccaa gttaaggagg gttgatttct gagaatagat gtgggcaaag ggcttcaga      840
cccagatggg gaattaggag atggcttggg aatgacatga tctgaggaag ggcctgggtc      900
aggggaagagc atgtttgtgt agttgggtca taacttctct gggcttctgc ttgtctaggt      960
gcaattaaca gactttgaga actctgccta tgtggtgttt g                                     1001

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&lt;210&gt; 178

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-354-72 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 10-354-72.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-354-72.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 430..447

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 830..849

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-354-72 potential probe

&lt;400&gt; 178

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tagcttttgt taaacacttg gctgttttgt tttcctaaaa gagcacactc gaagaaaatc      60
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ttccttaatt aaataccaga tcattctaat attttcttgg atgtgtgaaa agttgggttt      180
cttctctgct tttgtatcac agaatatcca gtccacaaat tggcagacaa tgagatttaa      240
gccccctcct ccaaactcag acattggatg gagagtagaa tttcgacca tggaggttaa      300

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149

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acacatgcat cagcaagaac tgactcaaaa gtactctttc gccagctgtt catcaccctt 360
ctgatgctgc tatgagaagg ctcttatata taaaataaaa attaatgatg tgcattggtg 420
caagtcagtc caagttaagg aggggttgatt tctgagaata gatgtgggca aagggtctcc 480
agaccagat ggggaattag ragatggctt ggggaatgaca tgatctgagg aagggtctgg 540
gtcaggggaag agcatgtttg ttaggttggt tcataacttc tctgggtctc tgcttgctta 600
gggtgcaatta acagactttg agaactctgc ctatgtgggtg tttgtgggtac tgctcaccag 660
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tcccacctaa tttcatttga aattgtttac cctttgcaat aaaatgtttc tccatcagcc 840
ctattcttta ttttctttta aacctaaagt taatttctag acgctttcca ttcaagatga 900
ttgatggcta gaagtagtcc tacacattga tttcatccac caaaaaatcc caggttgatt 960
acaatgttaa attaggatgg ggaggggaaag tgtaactgct g 1001

```

&lt;210&gt; 179

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-354-320 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-354-320.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-354-320.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 182..199

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 582..601

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-354-320 potential probe

&lt;400&gt; 179

```

ctccaaactc agacattgga tggagagtag aatttcgacc catggaggta agacacatgc 60
atcagcaaga actgactcaa aagtactctt tcgccagctg ttcacacccc ttctgatgct 120
gctatgagaa ggctcttata tataaaataa aaattaagta tgtgcatggt agcaagtcag 180
tccaagttaa ggagggttga tttctgagaa tagatgtggg caaagggtct ccagaccag 240
atggggaatt aggagatggc ttgggaatga catgatctga ggaagggtcct gggtcaggga 300
agagcatgtt tgtgtagtgt gttcataact tctctgggct tctgcttgct taggtgcaat 360
taacagactt tgagaactct gcctatgtgg tgtttgtggg actgctcacc agagtgatcc 420
tttcttacia attggatttt ctcatccac tgtcaaagggt aaggatatgt ttctttatgg 480
tgatgggtat agatctatct rtatatatat ttatttatat atgctattta tttcccacct 540
aatttcattt gaaattgttt accctttgca ataaaatgtt tctccatcag cctattctt 600
tattttcttt taaacctaag cttaatttct agacgcttct cattcaagat gattgatggc 660
tagaagtagt cctacacatt gatttcaccc accacaaaat cccagggttg ttacaatgtt 720
aaattaggat ggggagggaa agtctaactg ctggaatgcc ggtaaaagtc tctcagtag 780
aactgaactt gagatatttc tgaagaattc ggatgggatt gaactctcag aaaaacaaat 840

```

150

```

tggagttaat ctgcccccca aaagacccaaa aaacaagctt tctgggagga gtttgccagc   900
aaaacaactc caggtgatga gggagcagag ctgtgggttg tggagcagta gctgaggagg   960
tgaacgcctt tggtttcttg gccagggttc ctcttggcag a                               1001

```

&lt;210&gt; 180

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-354-360 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-354-360.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-354-360.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 142..159

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 542..561

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-354-360 potential probe

&lt;400&gt; 180

```

catggaggta agacacatgc atcagcaaga actgactcaa aagtactctt tcgccagctg   60
ttcatcacc ttctgatgct gctatgagaa ggctcttata tataaaataa aaattaagta   120
tgtgcatggt agcaagtcag tccaagttaa ggagggttga tttctgagaa tagatgtggg   180
caaagggtt ccagacccag atggggaatt aggagatggc ttgggaatga catgatctga   240
ggaagggcct gggtcaggga agagcatggt tgtgtagttg gttcataact tctctgggct   300
tctgcttgct taggtgcaat taacagactt tgagaactct gcctatgtgg tgtttgggt   360
actgctcacc agagtgatcc ttccctacaa attggatttt ctcatccac tgtcaaagg   420
aaggatatgt ttctttatgg tgatgggtat agatctatct gtagatata ttatttatat   480
atgctattta ttcccacct ratttcattt gaaattgttt accctttgca ataaaatggt   540
tctccatcag ccctattctt tattttcttt taaacctaa cttaatttct agacgtttc   600
cattcaagat gattgatggc tagaagtagt cctacacatt gatttcatcc accacaaaat   660
cccagggtga ttacaatggt aaattaggat ggggaggga agtgtaactg ctggaatgcc   720
ggtaaaaagt tcctcagtag aactgaactt gagatatttc tgaagaattc ggatgggatt   780
gaactctcag aaaaacaaat tggagttaat ctgcccccca aaagacccaaa aaacaagctt   840
tctgggagga gtttgccagc aaaacaactc caggtgatga gggagcagag ctgtgggttg   900
tggagcagta gctgaggagg tgaacgcctt tggtttcttg gccagggttc ctcttggcag   960
agagtggctg ggaaaaactg ctaaataag agggccctgc c                               1001

```

&lt;210&gt; 181

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

151

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<220>
<221> allele
<222> 501
<223> 10-355-87 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 10-355-87.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-355-87.mis2, potential complement

<220>
<221> primer_bind
<222> 415..434
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 821..840
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-355-87 potential probe

```

```

<400> 181
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ttgaaccagg gagacagagg ttgggggtgag atgagattgc accactgcac tccaacctgg      120
gcaacagagc gagactccat ctcaaaaaaa cagaagaaga actgggttct agtcctgcct      180
tctgccttcg ttaccctggg actcctgggtg agacactgct atataaaggg aaccagccac      240
ctgccctccc aacactgtat atctgttgca aggctcagtc acagagggtgc tttgaaatgc      300
agacgttcca catgaatata agttttgggt tttaagtttc tcttaaaatt ccctgattgg      360
ttgaaaaatc acataccaat ccatgattga gtgatacagt cctcacataa ataaaaagcc      420
tgttcctccc actgggaagc atttgagcag aaagtagaat ccctaaggga atgagtccta      480
attgcaaagg tgaggaaggt rctgcaactt gtatgtctga tcaagaatat tttgacttag      540
tcgtgattct ttttttcct taggttgatg agaacatgaa ggtagcacag aaaagagatg      600
ctgtcttgca gggaaatgtt tatttcagga aagatatttg caaaggattt acattatctt      660
agattttctaa tgtcagctta tgctgcaaca agctgctaaa gctccctga ccctcttctc      720
cggcaggtgg caatgcagtg gtggatggtt gtggcaaggc ccagaacagc acggagctcg      780
ctgcagagga gtacaccctc atgagcatag acaccatcat caatgggaag gtaggagggg      840
ggccctgtac agtacgcctc actgtgcaca tcccccgga gggagttcat aaagtgtcct      900
gcttcttgta ggaaggtgtg tttcctggac tgatcccaat tctgaactct taccttgaaa      960
acatggaagt ggatgtggac accagatgta gtattctgaa c                               1001

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```

<210> 182
<211> 1001
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele
<222> 501
<223> 10-358-60 : polymorphic base A or G

```

```

<220>
<221> misc_binding
<222> 481..500
<223> 10-358-60.mis1, potential

```

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-358-60.mis2, potential complement  
  
 <220>  
 <221> primer\_bind  
 <222> 442..460  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 870..889  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-358-60 potential probe

<400> 182  
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 caatttccca cttcattcac tctacttact gttctttatt ctctccacct ctttgatcaa 120  
 agatcgctca aggacctac agtcctctga ggcagtctca tgacactggc atttgtagtc 180  
 ctagccctct ggccaaatgg gagggcacat gttcttgtag acatgtgctg gtcctgttgc 240  
 cttaagagct ggcagtgtca gcatatgggt ggcattctaac aggtaccata gaaactggat 300  
 gtatgtgttc tcattgcctg cctgcaacct agaattcctg cccaggagga gttcttcaaa 360  
 cgctatgaaa gatgttccca ccccttgcca tgcagctaca ctaaaaggac atattgattt 420  
 ctctccagaa ttgtgtttat gctgaccact aaatatcaac ttattaaaaa aaaaacttac 480  
 gtggttttaa ttttttttcc rctccccct cgcccacagg agaactaatg acagttgccca 540  
 gatggatgag ggagtttatc gcaaaccatc ctgactacaa gcaagacagt gtcataactg 600  
 atgaaatgaa ttatagcctt attttgaagt gtaaccaaat tgcaaatgaa ttatgtgaat 660  
 gccagaggt acttgatca gcatttagga aagtaaaata tagtggaagt aaaactgact 720  
 catccaacta gacattctac agaaagaaaa atgcattatt gacgaactgg ctacagtacc 780  
 atgcctctca gccagcccggt gtgtataata tgaagaccaa atgatagaac tgtactgttt 840  
 tctgggccag tgagccagaa attgattaag gctttctttg gtaggtaaat ctagagttta 900  
 tacagtgtac atgtacatag taaagtattt ttgattaaca atgtatttta ataacatata 960  
 taaagtcatac atgaactggc ttgtacattt ttaaattctt a 1001

<210> 183  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens  
  
 <220>  
 <221> allele  
 <222> 501  
 <223> 12-468-63 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-468-63.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-468-63.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 439..458

153

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 946..966

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-468-63 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 376,386,618

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 183

```

cagtgc aaag acccagaacc cagccttcct tcacctgaaa cagctgcagc gtgcacacac      60
acttcttg tg catatatctc tgggtctattt attttcaaaa ctaagtataa tccaaattga      120
atgctggaa cagatttttc cattctgaat gtgtttaaat aacatggcca aggataattt      180
ctttctgtt tcttgacatt tataggtagt ataaaatatg tgacttccaa ataaacataa      240
accatcacac ttcacgaaaa aagagatctt tcgactttta actgcctgtt tcctatctta      300
attacaagta tttctaaaga aaaccattaa gttctagagc ttacagtaca tggtttttaa      360
attgtataag tgctgnttaa gagtanttga ttgccttttc ttggtttaat attaaaccag      420
ggtgttctgc caaagaccgc atagtttctc tgatatgggt ccaaagggat attccttcca      480
ctgaaacact gtaggtatac yagtaaatgc ttcctaattg tagatgataa atatttttgt      540
tgcttgtaga tagaattttt ccatctagtg taagcttagg attgttttct tttcccag      600
gacatgtaca gtttcacnta ctccacttaa aaaaaatcgt tagctcagat aaagtgtgtg      660
gcacatgaaa tgaattttgc caattcacca ccgaaatccc tgctttatat ctcttcagtg      720
gaacactaaa ccaacttctt tcccaagtac actgatttga tctttacaaa ttatatgaat      780
gtattaaatt atcacatgtg ccctgaaact gtgtacatct attatgtatc attagagaga      840
tggaaaaaaa aaaaagaaac tgcttagtaa agtagaaaaa atagttttaa agtaattttg      900
taaggcta at ggaagactta ctgtataaaa caaaaaggat ttaaccaca ttcaaattat      960
tgactgtttt ggggcttttc aggaatcact taaaaagcac c                                1001

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&lt;210&gt; 184

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-468-388 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-468-388.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-468-388.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 113..132

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

154

<222> 620..640  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-468-388 potential probe

<220>  
 <221> misc\_feature  
 <222> 50,60,292,676  
 <223> n=a, g, c or t

<400> 184  
 ttaagttcta gagcttacag tacatgggtt taaaattgta taagtgctgn ttaagagtan 60  
 ttgattgcct tttcttggtt taatattaaa ccagggtggt ctgccaaaga ccgcatagtt 120  
 tctctgatat gggtccaaag ggatattcct tccactgaaa cactgtatgt ataccagtaa 180  
 atgcttccta atggtagatg ataaatattt ttgttgcttg tagatagaat ttttccatct 240  
 agtgtaagct taggattggt ttctttttcc cagtgcacat tacagtttca cntactccac 300  
 ttataaaaaa tcggttagctc agataaaagt tgtggcacat gaaatgaatt ttgccaatc 360  
 accaccgaaa tccctgcttt atatctcttc agtggaaacac taaaccaact tctttcccaa 420  
 gtacactgat ttgatcttta caaattatat gaatgtatta aattatcaca tgtgccctga 480  
 aactgtgtac atctattatg yatcattaga gagatggaaa aaaaaaaaag aaactgctta 540  
 gtaaagtaga aaaaatagtt ttaaagtaat tttgtaaggc taatggaaga cttactgtat 600  
 aaaacaaaaa ggattttaac cacattcaaa ttattgactg ttttggggct tttcaggaat 660  
 cacttaaaaa gcaccnaagt tcacagccag gcacggtggc tcatgcctgt aatcccagca 720  
 ctttgggagg ctgaggtggg cagatcactt gaggtcagga gtttgagaca agcctggtca 780  
 tcatggcaaa atctcatctc tactaaaaat acaaaaatta gacctggtgg catgtgcctg 840  
 taatcccagc tactgaggag tctgaggcat gagaaacgct tgaacctggg aggcggaggt 900  
 tgcagtgagc caagatcgtg ccaactgcatt ccagcctggg gaacagagca agtctttgtc 960  
 tcaaaaaaaa aaaaaaaaaa agcactaagt tcatgacagg t 1001

<210> 185  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-468-491 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-468-491.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-468-491.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 10..29  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 517..537  
 <223> downstream amplification primer, complement

<220>

155

<221> misc\_binding  
 <222> 489..513  
 <223> 12-468-491 potential probe

<220>  
 <221> misc\_feature  
 <222> 189,573,901,905  
 <223> n=a, g, c or t

<400> 185  
 ccaaagaccg catagtttct ctgatatggt tccaaaggga tattctttcc actgaaacac 60  
 tgtatgtata ccagtaaatg cttcctaata gtagatgata aatatttttg ttgcttgtag 120  
 atagaatttt tccatctagt gtaagcttag gattgttttc tttttccag tgacatgtac 180  
 agtttcacnt actccactta aaaaaaatcg ttagctcaga taaagtgtgt ggcacatgaa 240  
 atgaattttg ccaattcacc accgaaatcc ctgctttata tctcttcagt ggaacactaa 300  
 accaacttct tcccaagta cactgatttg atctttacaa attatatgaa tgtattaaat 360  
 tatcacatgt gccctgaaac tgtgtacatc tattatgtat cattagagag atggaaaaaa 420  
 aaaaaagaaa ctgcttagta aagtagaaaa aatagtttta aagtaatttt gtaaggctaa 480  
 tggaagactt actgtataaa rcaaaaagga ttttaaccac attcaaatta ttgactgttt 540  
 tggggctttt caggaatcac ttaaaaagca ccnaagttca cagccaggca cgggtggctca 600  
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 tgagacaagc ctggtcacatc tggcaaaatc tcactcttac taaaaataca aaaattagac 720  
 ctggtggcat gtgcctgtaa tcccagctac tgaggagtct gaggcattgag aaacgcttga 780  
 acctgggagg cggaggttgc agtgagccaa gatcgtgcc ctgcattcca gcctggggaa 840  
 cagagcaagt ctttgtctca aaaaaaaaaa aaaaaaagc actaagttca tgacaggtta 900  
 nggttagccc ctcccacccc ccccaaagc catgtgactt caaatctggg ttgtagataa 960  
 agtttagatt agatgtttca gtttctaaca tatatgttcc t 1001

<210> 186  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-469-132 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-469-132.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-469-132.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 370..387  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 812..832  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-469-132 potential probe

156

```

<220>
<221> misc_feature
<222> 16,226,688,735
<223> n=a, g, c or t

<400> 186
gcccgcggtc cccgtncccc gccgggcacc cgtgtctggc cgcccgcgct gcctttgtct      60
tgccctacgc ccccggccct atcgcggtcc gcgttctcca ctgcgcccgg gggtcctcga      120
gcgcctcccg gcccgggag catcgtcctt gctttcgccct agtcgcgctc cagggccatc      180
ctcctgacta gactcctttc ttgttcattc attcttttgg gttttneccc ttcattccat      240
aaacacggat tgggcgctgg gagggcgatga gcatgaaaaa gtgacattag gtgctgcgag      300
tcttcaacct agtgtatctg cagaccggca actaccatga tccaccttac aggtcaaaaa      360
ttcgcttttg cgggtcaatat gaaagagatt taagttttcc tagtaggcct gctgctttta      420
gaaatcaggt cttcaggatg tccacttctg ccctacctca tgcagggtgc ttttccttaa      480
acctctccaa cttctcatct yccctcccct ccaaaacaca cacacttcat agctataaat      540
tgaaaccagt aaattacggc tcagtttcct acatggggga ggaaccttcc ctttcctggt      600
taattctaag tgggaatgca tttagagttt caacattagg atttgcttgt gtctttccat      660
ggcctaggtc cagcaactaa actttacnaa accagtgttc tgcctagaga tgaaatgtct      720
ctagattctc agtanggttg aaccattggt tattggaatg atctggaatg cagcaatggt      780
aaaactgttg atttagtgct atctgccaaa actttttgtg ccttcccctt ctgtttttga      840
aaattacgtg tgtatgcagt acacattaaa tgacttcttg agaagtccat ctctgttttt      900
gttttttagct tttgccttag atggcttgcc acctgtgtaa attgtttatc ttttggccag      960
tgttgttatc aatcttgcc taagtgcga cgtgctgtta c                                1001

<210> 187
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-469-245 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-469-245.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-469-245.mis2, potential complement

<220>
<221> primer_bind
<222> 257..274
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 699..719
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-469-245 potential probe

<220>
<221> misc_feature
<222> 113,575,622,936
<223> n=a, g, c or t

```



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<400> 187
tcctcgagcg cctcccggcc ccgggagcat cgtccctgct ttgcgcctagt ccgcgtccag    60
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attccataaa cacggattgg gcgctgggag ggcgtgagca tgaaaaagt acattagggtg    180
ctgcgagtct tcaacctagt gtatctgcag accggcaact accatgatcc accttacagg    240
ctcaaaattc gcttttgcg tcaatatgaa agagatttaa gttttcctag taggcctgct    300
gcttttagaa atcaggctct caggatgtcc acttctgccc tacctcatgc aggtgtcttt    360
tccttaaacc tctccaactt ctcattcttc ctcctctcca aaacacacac acttcatagc    420
tataaattga aaccagtaaa ttacggctca gtttctaca tgggggagga accttccctt    480
tcctgtttaa ttctaagtgg raatgcattt agagtttcaa cattaggatt tgcttggtgc    540
tttccatggc ctaggtccag caactaaact ttacnaaacc agtggtctgc ctagagatga    600
aatgtctcta gattctcagt anggttgaa cattgtttat tggaaatgatc tggaaatgcag    660
caatgttaaa actgttgatt tagtgtcatc tgccaaaact ttttgtgctt tccctttctg    720
tttttgaaaa ttacgtgtgt atgcagtaca cattaaatga cttcttgaga agtccatctc    780
tgtttttgtt tttagctttt gccttagatg gcttgccacc tgtgtaaatt gtttatcttt    840
tggccagtgt tgttatcaat cttggcctaa gtgtgacgt gctgttacac cagtgcagaa    900
tatctccttt caaaaagtta gttttttttt tttttnctaa gattgggatt ctacctcta    960
cagctgaaga aattgtgctg ttttaaatta ggaaagtatg a                                1001

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<210> 188

<211> 1001

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 501

<223> 12-472-435 : polymorphic base A or G

<220>

<221> misc\_binding

<222> 481..500

<223> 12-472-435.mis1, potential

<220>

<221> misc\_binding

<222> 502..521

<223> 12-472-435.mis2, potential complement

<220>

<221> primer\_bind

<222> 68..86

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 533..553

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 12-472-435 potential probe

<400> 188

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gatttggttaa atgtgaagtt tgaatgtggc cattttccct cccccatact tcatgtcctc    60
acattagaga gaggatgatt taagtgaatc aaaccceaagg aactggattc ttcttggtat    120
atttcactgt aagataggca caggtagatg tgctctgtat ggggtgcata aacatgcttt    180
ttgatcagaa atataactgc atggagcttt ttttagcatg taagtgcacac tttgaatttg    240
caggagctga cttttgtttg tttttagatg gctttgagtg ctgcattctc cttttaccga    300
ggctatgtgt cagacattga ttgtcgctgg ggagtgattt ctgcattctg agatgataga    360
actcgggagg agcgaggact ggaggtggga attgtttttc cttaatagcc cttttaagtc    420

```

158

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aagcaggtaa aatggatctt ttgtaactac ttgcaatttc aggatgtctc cctgcaattt 480
ttatctgaaa tggggaaaaa ratttggtct aggtggggag ttaattttta tcgtatgttg 540
atgtctgtat ttgttttagc ttcattttaa attagtagtc cgatttttct atattttgga 600
agtgtgatg gctgttttta ccaaaaggct ttttggtgcc acatgagtgt tttgtatatg 660
gtgctttctg ctgtggaaag tataatttgt tgtcagagat tctttttcca tgtaaaagg 720
taggcattga ctaataaggg tgagatggca cctcttgttt gcatgttgaa ctgtaaagta 780
acccttgcat ctctccaaca aggggtgtgc atcagaacca cccatggaaa tctttgaaaa 840
tagacacca gaccctatcc tagacctact aaataagaat ctctaggatg ggacccgaac 900
aagggtgttt ttaaaggagc cctcataagt ggttttgata ttctcaacca gggctatact 960
actactgctc tggctcttcat gtaattgttc ataaattgtt g 1001

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&lt;210&gt; 189

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-473-311 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-473-311.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-473-311.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 192..210

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 740..758

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-473-311 potential probe

&lt;400&gt; 189

```

taatttttat cgtatgttga tgtctgtatt tgttttagct tcatttaaaa ttagtagtcc 60
gattttttcta tatttttgaa gtgctgatgg ctgtttttac caaaaggctt tttggtgcc 120
catgagtgtt ttgtatatgg tgctttctgc tgtggaaagt ataatttggt gtcagagatt 180
ctttttccat gtaaaagggt aggcatggac taataagggt gagatggcac ctcttgtttg 240
catgttgaa cgtaaaagtaa cccttgcatc tctccaacaa ggggtgtgca tcagaaccac 300
cctaggaaat ctttgaaaat agaccccag accctatcct agacctacta aataagaatc 360
tctaggatgg gaccgaaaca aggggtgttt taaaggagcc ctcataagtg gttttgatat 420
tctcaaccag ggctatacta ctactgctct ggtcttcatt taattgttca taaattgttg 480
aacatgcttt ctaagtagaa mtaaaactta aaggataatt attttgatag gtttatctta 540
cgttatagat ttatttatat atttgctttt aaccttagta tttattttcc ctagcttaat 600
aagaatgaaa gtgatctgct tttaaaagat aaaagtatat tttaatcata ccagaaaaaa 660
agggtattta aattgccttc agatttttaa aaacataatt ttccttaata atacttttta 720
gggtcacta actttttttc tgtatcttat gttgaggttt ttatataatt atcatatata 780
aaaatatatt ccactcattt tatatagcat attcatttat atatatattt aatcttttagc 840
cattgaagaa caataactat aggatcagta aatcccagata tgactcaata gacagctatt 900
tatctaagtg tggtgagaaa tataatgaca tcgacttgac gatagataaa gagatctacg 960

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159

aacagctgtt gcaggaaggt gggtttctac tccatctttc t 1001

<210> 190  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-473-483 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-473-483.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-473-483.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 20..38  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 568..586  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-473-483 potential probe

<400> 190  
 cagagattct tttccatgt aaaagggttag gcattgacta ataaggggtga gatggcacct 60  
 cttgttttga tgttgaactg taaagtaacc cttgcatctc tccaacaagg ggtgtgcatc 120  
 agaaccaccc atggaaatct ttgaaaatag acaccagac cctatcctag acctactaaa 180  
 taagaatctc taggatggga cccgaacaag ggtgttttta aaggagccct cataagtgg 240  
 tttgatattc tcaaccaggg ctatactact actgctctgg tcttcatgta attgttcata 300  
 aattgttgaa catgctttct aagtagaaat aaactttaaa ggataattat tttgataggt 360  
 ttatcttacg ttatagattt atttatttat ttgcttttaa ccttagtatt tattttccct 420  
 agcttaataa gaatgaaagt gatctgcttt taaaagataa aagtatattt aaatcatacc 480  
 agaaaaaaag ggtattttaaa ytgcttcag attttaaaaa acataatttt ccttaataat 540  
 acttttttag gctcactaac tttttttctg tatcttatgt tgagggtttt atataattat 600  
 catatataaa aatatattcc actcatttta tatagcatat tcattttatat atatttttaa 660  
 tcttttagcca ttgaagaaca ataactatag gatcagtaaa tcccgatatg actcaataga 720  
 cagctattta tctaagtgtg gtgagaaata taatgacatc gacttgacga tagataaaga 780  
 gatctacgaa cagctgttgc aggaagggtg gtttctactc catctttctg ggtttgaatg 840  
 tacgccttta gttcttcaaa gctcttttac acttttttgc tgactcctgg tctgggttct 900  
 attttttagt caaagtcttt aacttctctc atgaggctac ttgttcctaa agtttcaggt 960  
 tccaaatact tgtgagattt tcttgatttt tagcaaaagg a 1001

<210> 191  
 <211> 998  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele

160

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<222> 501
<223> 12-475-85 : polymorphic base T or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-475-85.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-475-85.mis2, potential complement

<220>
<221> primer_bind
<222> 566..585
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 108..126
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-475-85 potential probe

<220>
<221> misc_feature
<222> 555,666,713
<223> n=a, g, c or t

<400> 191
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taatgtgact gcagccttcc cagctaattct gctggcaagc gagccctgtg aaaaggcaga      120
tgcaggacga gcccttttcta tgaaaatcag gctgacctca tttgatagtt atgcagcctg      180
tgtcgatctc ttcccagggc agcaggtagc aggccagaat cagaacccca agtcaggagc      240
tttaacttat ctgagatttt catccttttag catgaatttc aaatatggca gggaaggaga      300
ttgaattcac aaaaacctaa atcttccaaa tcccacaggt tacagttact ctgttggtgg      360
gatccactat aacacccagg tacgattcct gaatgggaac ctgatgctgc ctttgtgaga      420
tgtggttatg acacagtgtc tgcactcaga ttcgtatgat ggtggaatgg cattaactt      480
ggaggtcttg aagccctgtc ktcctgtccc tggaggcata ggtgaccccc agtgagcact      540
agcagtcact ttggnactta gaattcctgt atttagcatt gactctcact tagtctcttt      600
agaaaaactct cttttttctc tctgttaggt tgattagtct aacttcagtt cctaggaact      660
aaaatnccctg aattttctcat aggcagtctg aatttagggc agcctaattt atnaacatgc      720
cttagaagga aatcatgtgt gcattcttgc ttgctcagcc caagaagtgt atcaggccct      780
gagcggggca ccgcccactg ggggatgtgt tctctgtctt gtcccatgc taaagcccta      840
aactctctctg cacctagagg gaaccgctgc agaaattctc catgattctc acccactaca      900
gtgccccagc tctggctact gcagggcagt tgatccaaag cagcttctgt ctggggatca      960
ggatactgtt tcttgtcaac cacgtggtga ctagaact      998

<210> 192
<211> 985
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 485
<223> 12-475-446 : polymorphic base G or A

<220>

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161

<221> misc\_binding  
 <222> 465..484  
 <223> 12-475-446.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 486..505  
 <223> 12-475-446.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 911..930  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 453..471  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 473..497  
 <223> 12-475-446 potential probe

<220>  
 <221> misc\_feature  
 <222> 47,196,202..203,900  
 <223> n=a, g, c or t

<400> 192  
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 gcctgcactc ttagctgctc cccacacctat taccgcccac ataggcagga ggaaaagagc 120  
 tctgtgacag ttttaaatagg ctccaaacag agatgagatt gaggggtggcg aggaagagaa 180  
 gaaagaatat agaaanggga tnnnggtggta atgcaggaag taggcttttt gttttttaat 240  
 gaaaatatga atatgtcatc agccacagat ccacgcttaa gcattttttg gtaccccata 300  
 gcaggttaatt ctcttctagt gaaatgccag cactctttca aaacgacacc attgaatatg 360  
 cttggtaaaa ttctcgtctc gcaggacaaa tgctaacaag tttagtaatg tgactgcagc 420  
 cttcccagct aatctgctgg caagcgagcc ctgtgaaaag gcagatgcag gacgagccct 480  
 ttctrtgaaa atcaggctga cctcatttga tagttatgca gcctgtgtcg atctcttccc 540  
 agggcagcag gtagcaggcc agaatcagaa ccccaagtca ggagctttaa cttatctgag 600  
 attttcatcc ttagcatga atttcaaata tggcagggaa ggagattgaa ttcacaaaaa 660  
 cctaaatctt ccaaattcca caggttacag ttactctgtt ggtgggatcc actataacac 720  
 ccaggtacga ttctgaatg ggaacctgat gctgcctttg tgagatgtgg ttatgacaca 780  
 gtgtctgcac tcagattcgt atgatgttg aatggcatta aacttggagg tcttgaagcc 840  
 ctgtcgtcct gtccctggag gcataaggta cccccagtga gcactagcag tcactttggg 900  
 acttagaatt cctgtattta gcattgactc tcacttagtc tctttagaaa actctctttt 960  
 ttctctctgt taggttgatt agtct 985

<210> 193  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-477-100 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-477-100.mis1, potential

162

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<220>
<221> misc_binding
<222> 502..521
<223> 12-477-100.mis2, potential complement

<220>
<221> primer_bind
<222> 580..600
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 62..82
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-477-100 potential probe

<220>
<221> misc_feature
<222> 368
<223> n=a, g, c or t

<400> 193
tcagttactg tgacctacca tatcatgggt gaattcctcc atggttaagac cctactgtct 60
agaagtatgg tggtagcaga tgtactcatt gattgggtact gtctgtagtt tcattttgaa 120
gcataaacct gagtaacatt cagtaagact ctgtatgttc agaaaagatg tcacaatatt 180
aattgagcct ctaacatcac tatttatgtt tctgaacatt ttacatact gtcaaggcat 240
ctgaatatat tcttaaagtt agaaaactgt atatgaagca ttactgatat tgctgggtatt 300
actggaaaga ccatatccta acgagcttgc taattagtac tctgtttcta catttcatga 360
tataatcnag gtgtttttaa aaaaacattt taacagcttt attgagttat aattaacata 420
aagtttaactg cacatattta aaatgtacaa tttggttaagt ttgatatat gtatacatca 480
taaaatcacc tctgataata rgtatattca ttatctccaa aaatgtcatc atgccctttt 540
ggtaatttct tcctctggcc atcccatgca gctgatttgc tttctgtagt ttgcattttc 600
tagattttca tatttaagaa atcatagttt gcatttgata tctagtttca tatacctagg 660
aaatcatcca gcataccttc ttttggtctg ctttcttaga gtctacataa atattttgag 720
atccatccat gttgttacat gtgtcagtag ttcatctcct ttcgtcgtcg agtaatat 780
cattccatgg ttataccagt ttgtttatac attcaccatt tgtgttgcat tcagtttttc 840
actattataa aaaaattata tgaacattaa catacaagtt tgtatgtatg gactaatgct 900
tttcttcac ttattaccct aggagtagcg tgttggttaac tttttaagga aactataata 960
ccaaactttt ccagggtggt tataccattt tgcattctca c 1001

<210> 194
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-477-331 : polymorphic base G or A

<220>
<221> misc_binding
<222> 481..500
<223> 12-477-331.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-477-331.mis2, potential complement

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<220>
<221> primer_bind
<222> 812..832
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 294..314
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-477-331 potential probe

<220>
<221> misc_feature
<222> 600
<223> n=a, g, c or t

<400> 194
gtgtaacaaa ggaggaaagg tgagaaatac cattggatag tatgtgaagg ttattgtgca      60
gttttagtgt gtaaatgcag tgaatagggt tagccagctt gtagaagtct ttgtctagtt      120
ctcaaaaagt agagggttat ttgtacttta aaagcatgag aaaagtagaa tcgaaaaaaaaa      180
gtgtattagt cttctatacc tatctcctac agaaaaaatc ttttaagact tatcagttac      240
tgtgacctac catatcatgg ttgaattcct ccatgttaag accctactgt ctagaagtat      300
ggtgttagca gatgtactca ttgattggta ctgctgtag tttcattttg aagcataaac      360
ctgagtaaca ttcagtaaga ctctgtatgt tcagaaaaga tgtcacaata ttaattgagc      420
ctctaaccatc actatttatg tttctgaaca tttttacata ctgtcaaggc atctgaatat      480
attcttaaag ttagaaaact rtatatgaag cattactgat attgctggta ttactggaaa      540
gaccatatcc taacgagctt gctaattagt actctgtttc tacatttcat gatataatcn      600
agggtgtttt aaaaaaacat tttaacagct ttattgagtt ataattaaca taaagttaac      660
tgacacatatt taaaatgtac aatttggtta gttttgatat atgtatacat cataaaatca      720
cctctgataa taagtatatc cattatctcc aaaaatgtca tcatgccctt ttggtaattt      780
cttcctctgg ccatcccatg cagctgattt gctttctgta gtttgcattt tctagatttt      840
catatttaag aaatcatagt ttgcatttga tatctagttt catataccta ggaaatcatc      900
cagcatatcc tcttttggtc tgccttctta gagtctacat aaatattttg agatccatcc      960
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<210> 195
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-477-332 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-477-332.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-477-332.mis2, potential complement

<220>
<221> primer_bind
<222> 813..833

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164

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<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 295..315
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-477-332 potential probe

<220>
<221> misc_feature
<222> 601
<223> n=a, g, c or t

<400> 195
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agtttagtgt tgtaaatgca gtgaataggt ttagccagct tgtagaagtc tttgtctagt      120
tctcaaaaag tagaggttat gttgtacttt aaaagcatga gaaaagtaga atcgaaaaaa      180
agtgtattag tcttctatac ctatctccta cagaaaaaat cttttaagac ttatcagtta      240
ctgtgaccta ccatatcatg gttgaattcc tccatgttaa gaccctactg tctagaagta      300
tggtgttagc agatgtactc attgattggt actgtctgta gtttcatttt gaagcataaa      360
cctgagtaac attcagtaag actctgtatg ttcagaaaag atgtcacaaat attaattgag      420
cctctaacat cactatttat gtttctgaac atttttacat actgtcaagg catctgaata      480
tattcttaaa gttagaaaac ygtatatgaa gcattactga tattgctggt attactggaa      540
agaccatatc ctaacgagct tgctaattag tactctgttt ctacatttca tgatataatc      600
naggtgtttt taaaaaaaaca ttttaacagc tttattgagt tataattaac ataaagttaa      660
ctgcacatat ttaaaatgta caatttggtg agttttgata tatgtataca tcataaaatc      720
acctctgata ataagtatat tcattatctc caaaaatgtc atcatgcccc tttggtaatt      780
tcttctctcg gccatcccat gcagctgatt tgctttctgt agtttgcatt ttctagattt      840
tcatatttaa gaaatcatag tttgcatttg atatctagtt tcatatacct aggaaatcat      900
ccagcatatc ctcttttggt ctgccttctt agagtctaca taaatatttt gagatccatc      960
catgttggtg catgtgtcag tagttcattc cttttcgtcg c                                1001

<210> 196
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-477-44 : polymorphic base C or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-477-44.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-477-44.mis2, potential complement

<220>
<221> primer_bind
<222> 524..544
<223> upstream amplification primer, complement

<220>
<221> primer_bind

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165

<222> 6..26  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-477-44 potential probe

<220>  
 <221> misc\_feature  
 <222> 312  
 <223> n=a, g, c or t

<400> 196  
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 tgaagcataa acctgagtaa cattcagtaa gactctgtat gttcagaaaa gatgtcacia 120  
 tattaattga gcctctaaca tcactattta tgtttctgaa catttttaca tactgtcaaag 180  
 gcatctgaat atattcttaa agttagaaaa ctgtatatga agcattactg atattgctgg 240  
 tattactgga aagaccatat cctaacgagc ttgctaatta gtactctgtt tctacatttc 300  
 atgatataat cnaggtgttt ttaaaaaaac attttaacag ctttattgag ttataattaa 360  
 cataaagtta actgcacata tttaaaatgt acaatttggg aagttttgat atatgtatac 420  
 atcataaaat cacctctgat aataagtata ttcattatct ccaaaaatgt catcatgccc 480  
 ctttggtaat ttcttctctt sgccatccca tgcagctgat ttgctttctg tagtttgcat 540  
 tttctagatt ttcataattta agaaatcata gtttgcatct gatattctagt ttcataatac 600  
 taggaaatca tccagcatat cctcttttgg tctgccttct tagagtctac ataaatattt 660  
 tgagatccat ccatgttggt acatgtgtca gtagttcatt ccttttcgtc gctgagtaat 720  
 atttcattcc atggttatac cagtttggtt atacattcac catttggtgt gcattcagtt 780  
 tttcactatt ataaaaaaat tatatgaaca ttaacataca agtttgatg tatggactaa 840  
 tgcttttctt catcttatta ccctaggagt agcgtgttgt taacttttta aggaaactat 900  
 aataccaaac ttttccaggg tgggtatacc attttgcatt ctcactagca gagtatgcca 960  
 gttcagttca tccacattct tgtaaacacc ggtatgggca g 1001

<210> 197  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 12-478-223 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-478-223.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-478-223.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 704..723  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 234..254  
 <223> downstream amplification primer

<220>

166

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<221> misc_binding
<222> 489..513
<223> 12-478-223 potential probe

<220>
<221> misc_feature
<222> 35,194,272,317,458
<223> n=a, g, c or t

<400> 197
tcttgctctg tcacccaggc tagagtgcag tgacnaccat cacagtgtgc tgcagccgca      60
gcctcccagg ctcaagctgt ttagatcgta tttctaagtt gcgcttcaaa aacgcattga      120
cacaggctca ggctaggtct cttaaaataa gatatagagc acaaattaat tatttcttaa      180
taggcattac aaangattca ggttccttct ataggtcata gcgtattctg taacacaaaa      240
agcaaagttg tcgcatctct ctatcttttt tnagctgtca ctggacattg attcaacagc      300
atttattgtc cgtaanccc ctaccacccc ccagaagttc ccttggtgcc tttgcacgtg      360
tgtgcatgcy tgtatgctag agagagaggg ggagagcact tcagttgatt ctgggtttcc      420
ttcagacacc acctttgaga aactgcctta aagagtgnct tggtcattaa ttagatgaag      480
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taccattgga tagtatgtga aggttattgt gcagtttagt gttgtaaatg cagtgaatag      600
gtttagccag cttgtagaag tctttgtcta gttctcaaaa agtagagggt atgttgtact      660
ttaaaagcat gagaaaagta gaatcgaaaa aaagtgtatt agtcttctat acctatctcc      720
tacagaaaaa atcttttaag acttatcagt tactgtgacc taccatatca tggttgaatt      780
cctccatgtt aagaccctac tgtctagaag tatggtgtta gcagatgtac tcattgattg      840
gtactgtctg tagtttcatt ttgaagcata aacctgagta acattcagta agactctgta      900
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<210> 198
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-478-320 : polymorphic base G or A

<220>
<221> misc_binding
<222> 481..500
<223> 12-478-320.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-478-320.mis2, potential complement

<220>
<221> primer_bind
<222> 801..820
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 331..351
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-478-320 potential probe
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167

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<220>
<221> misc_feature
<222> 132,291,369,414,555
<223> n=a, g, c or t

<400> 198
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tttattttatt tattttattta ttttgagaca ggggtctttct tgctctgtca cccagggtag      120
agtgcagtga cnaccatcac agtgtgctgc agccgcagcc tcccaggctc aagctgttta      180
gatcgtattt ctaagttgcg cttcaaaaac gcattgacac aggttcaggc taggcttctt      240
aaaataagat atagagcaca aattaattat ttcttaatag gcattacaaa ngattcaggt      300
tccttctata ggtcatagcg tattctgtaa cacaaaaagc aaagttgtcg catctctcta      360
tcttttttna gctgtcactg gacattgatt caacagcatt tattgtccgt taancccccta      420
cccacccccca gaagttccct tgtgcccttt gcacgtgtgt gcatgcgtgt atgctagaga      480
gagagggggga gagcacttca rttgattctg gttttccttc agacaccacc tttgagaaac      540
tgccttaaag agtgntgtgg tcattaatta gatgaagggg agaaaggaag gcattatggt      600
caaaagaaca gtgtaacaaa ggaggaaagg tgagaaatac cattggatag tatgtgaagg      660
ttattgtgca gtttagtggt gtaaatagcag tgaatagggt tagccagctt gtagaagtct      720
ttgtctagtt ctcaaaaagt agagggttatg ttgtacttta aaagcatgag aaaagtagaa      780
tcgaaaaaaaa gtgtattagt cttctatacc tatctcctac agaaaaaatc ttttaagact      840
tatcagttac tgtgacctac catatcatgg ttgaattcct ccatgttaag accctactgt      900
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<210> 199
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-479-289 : polymorphic base G or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-479-289.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-479-289.mis2, potential complement

<220>
<221> primer_bind
<222> 213..230
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 678..698
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-479-289 potential probe

<220>
<221> misc_feature
<222> 444,957..958
<223> n=a, g, c or t

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<400> 199
ctctcctggt actccccctt ctctcactcc actccagcca tactagcctc cttactgttc      60
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ggaatggctg aatttcctta cctccttcaa atccttgctt aatccctcct tctcaaagag      180
acctaaacac cctggccacc ttctttaata taatattctg accatgcacc taacctcaca      240
cctcgatccc ctgctctaata ctgtcttttt cataatgttt atcaacttct gactttctat      300
atagttatatt attgtgcatg ttgtgtgect ccccttgcta aaatgtaagc tccacaaggg      360
caggcatgct tgtttgttat agtcagtgat gtatcctaag cacctagaac agtacttgcc      420
tgatgggtgag tgctcaatga acantttgca gaaaaatgaa tgaatgaatg ccaggcattc      480
ataagcccac aaaaaaaatc katgtgtcat tctacacctt cactgtagcc taatgcttca      540
tgctccaaaa agaaagaaga ggcattcattt tctctgtttc ctgtgctcca cctctcgcac      600
attccccact ctctagctg ctgtctttcc cctgactttt ttgtctagta acaagaactc      660
cctgcctggg gaaagtcggc attttcttct caatttactt ctaaataatta gtgcttttgt      720
ttaccctgtg aagcagatgg taacagttaa tgtaataacc agactgtctc ctctctcttg      780
tttgacacag tagatgtcat gagccatatt ttcccttggg tatatttgtt aaatagaaga      840
caggagatct gctatatggt caacattagg tctggtcact gcttttatat tgtaaacact      900
gtgtttcctt tcatatatat acatcatttg gaaacatttg ccctagtcc agcaatnnag      960
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<210> 200
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-482-237 : polymorphic base G or A

<220>
<221> misc_binding
<222> 481..500
<223> 12-482-237.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-482-237.mis2, potential complement

<220>
<221> primer_bind
<222> 720..737
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 223..243
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-482-237 potential probe

<220>
<221> misc_feature
<222> 552,561,757,831,845
<223> n=a, g, c or t

<400> 200
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gagacaatat tgagaaagtt actccaccta atcctatagg aaattaattg atttttgtgc      120

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169

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agtattaaaa ttgagattaa tgtgttataa ataaaaagca gcctgctctg tgctacagaa 180
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atcatgtgaa attcttttagt atgaatagac atttcaaagt aggaggaaaa aaacctattt 300
tctcaaaata tatgaaatgt tagcagaatt tgatacagct ttagacaaaa ctgctaaaaac 360
attttctttt tggaacatag aacaatttaa ttattacagt atccacagtc atcaaatata 420
tatatattta aatcatcaaa tatatatggt taaaatatag ttgtgtgtgt gatgttatgt 480
atgtataatg tcatgtacac raaagcacat acagtttcta ctttgtactt actgggtccc 540
taaaagtgtc tnggagatta nccatgtcaa acagtggttt tcctatttgt gatgtcatca 600
ttagacatat ctaacatggg agtccatctt taaaaatctg agccatatgt aattttatta 660
catctgtatt agaatctcat attcagtttt ctacttttta aaacacagct cctcttttag 720
tcttgtcagt caggttgtcc tatgttactg tgggtgnttt aaaaattaag tagcatcact 780
atggctaaca cctctttggg tgctccaagt ttgggaagtg aatgccatcc naacttgatg 840
gttgnagatt tcttttaaaa aaaaatttgt cattaccatt ttgctgtgag aggaaacagt 900
cttctgggca gaaggcattg agtccctaac tttaaaacag ttataacct agaggggtgac 960
tacttggcat gatctctgtg ggggtggatgc agtgatctgc g 1001

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&lt;210&gt; 201

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-482-285 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-482-285.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-482-285.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 768..785

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 271..291

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-482-285 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 600,609,805,879,893

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 201

```

cccgcaactc tcaactgtctt tctctatttt attaaagtct tgaaattttt ctcttggggac 60
tcatatcact aatctaggca agttaagaat gttttctctg tttgggctga gacaatattg 120
agaaagttac tccacctaat cctataggaa attaatgtat ttttgtgcag tattaaaatt 180
gagattaatg tgttataaat aaaaagcagc ctgctctgtg ctacagaaaa ctagatcagg 240
gagaggccac atttatttta atttctaaaa gtgtaaacct ttcatagtat catgtgaaat 300
tcttttagtat gaatagacat ttcaaagtag gagggaaaaa acctattttc tcaaaatata 360

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170

tgaaatgtta	gcagaatttg	atacagcttt	agacaaaact	gctaaaacat	tttctttttg	420
gaacatagaa	caattttaatt	attacagtat	ccacagtcac	caaatatata	tatatTTTaaa	480
tcatcaaata	tatatgttta	waatatagtt	gtgtgtgtga	tgttatgtat	gtataatgtc	540
atgtacacaa	aagcacatac	agtttctact	ttgtacttac	tggtccccta	aaagtgcctn	600
ggagattanc	catgtcaaac	agtggttttc	ctattttgtga	tgatcatcatt	agacatatct	660
aacatgggag	tccatcttta	aaaatctgag	ccatatgtaa	ttttattaca	tctgtattag	720
aatctcatat	tcagttttct	acttttttaa	acacagctcc	tcttttagtc	ttgtcagtca	780
ggttgctcta	tgttactgtg	ggtgntttaa	aaattaagta	gcatcactat	ggctaacacc	840
tctttgggtg	ctccaagttt	gggaagtga	tgccatccna	acttgatggg	tgtagatttc	900
ttttaaaaaa	aaatttgta	ttaccatttt	gctgtgagag	gaaacagtct	tctgggcaga	960
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&lt;210&gt; 202

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-482-482 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-482-482.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-482-482.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 965..982

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 468..488

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-482-482 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 67,797,806

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 202

tctgcagtaa	gagaaatggt	ctactctata	ctatccatta	gggagtagga	aactgatttt	60
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aatggacaac	acaatttttt	gtgttccttt	ggtgacaggt	gacattttct	ccctgtatga	180
gagtcctccc	agttttacccc	gcaactctca	ctgtctttct	ctattttatt	aaagtcttga	240
aattttttctc	ttgggactca	tatcactaat	ctaggcaagt	taagaatggt	ttctctgttt	300
gggctgagac	aatattgaga	aagttactcc	acctaatect	ataggaaatt	aattgatttt	360
tggtcgagat	taaaattgag	attaatgtgt	tataaataaa	aagcagcctg	ctctgtgcta	420
cagaaaaacta	gatcagggag	aggccacatt	tatttttaatt	tctaaaagtg	taaacctttc	480
atagtatcat	gtgaaattct	wtagtatgaa	tagacatttc	aaagtaggag	gaaaaaaacc	540
tattttctca	aaatatatga	aatgttagca	gaatttgata	cagctttaga	caaaactgct	600

171

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aaaacatttt ctttttggaa catagaacaa ttttaattatt acagtatcca cagtcatcaa 660
atatatatat atttaaatca tcaaataatat atgtttaaaa tatagttggtg tgtgtgatgt 720
tatgtatgta taatgtcatg tacacaaaag cacatacagt ttctactttg tacttactgg 780
tcccctaaaa gtgcttngga gattanccat gtcaaacagt ggttttccta tttgtgatgt 840
catcattaga catatctaac atgggagtc atctttaaaa atctgagcca tatgtaattt 900
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tttagtcttg tcagtcaggt tgcctatgt tactgtgggt g 1001

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&lt;210&gt; 203

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-483-322 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-483-322.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-483-322.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 802..820

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 311..331

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-483-322 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 253,301..302,345,368,400,565,570,968,973,978

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 203

```

agacactttg aaatgtcttt gcttaaaaac aattggttta agtgactgc aaaagcatta 60
catgggtctag cctcataata atttccctt tttggagacc caggattcag tgtgggctct 120
gccagattt cagagatcta ggcaaaaaag aaataatccc tatatgaata aaattgggtct 180
cctcatacaa tcccatgata gagttctata attttatgtt tgatttggca tccatcttta 240
tttccctct agncaccact agactttttc tgtctgtacc ttgagatata aattttgcta 300
nntgattttt catctaagag ttgtttcctt caatatgcag gtttnagggc tatttagctg 360
acaactgncc aggttaatga aacaggttat catgagtttn gcaagtctaa gacaggggaa 420
aacaaaagga ggtcttagga atctataaga tgtacttcta tcagtatgcc taatacatct 480
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attggcttgc agaaaaataa aggtngcttn aaatccaatg ctttatcaga gaaaagaaaa 600
gactagccaa atgctttttc aagtttatgt gatttaagta aaatctttaa taaataagct 660
agctttaaaa ttactggcaa agtaatatta gaaatgtctt aagaattgcc agcatacatt 720
ttcgtttgca tttatggatc aagtcatttc atatttatcc ctgccaaata ctgtaagggt 780
tcaaagtttg gcataggggtt acaaaactat aaaccagcc taaaacagaa tgatttttgt 840

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172

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ttgtgttaat ttttaataaa taagacattg atattgggtt aatgaaaata gctacatctt   900
gaattattta gtaaaattac tgtaacttct aatcttgtgg ccttagggag tctagtccac   960
aggcaatnaa ggnnttcntt ttgggaaatg actgttatca t                        1001

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&lt;210&gt; 204

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-484-46 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-484-46.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-484-46.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 528..546

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 86..106

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-484-46 potential probe

&lt;400&gt; 204

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tgtgagcaag aagtgtgggg agtggttgta aaggggcaca gtatttacac aacatctgtt   60
ttgtaaaccac aaagccctct gaaaagggga tactagaaat gaggtgtctt tggcaggtgc  120
agtgatggca agactgcaca caattaattc ttgaaaacaa atcccatcta tttagttctt  180
caatttggtg tagattaggg tgctctgggt tcacactgcc cgaggtcaaa cccaaccag  240
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gcttcagggtg agtatggctt tgtgttgaaa tgcagatgcc cttcactaga cccctgactt  360
ctgagagtat agtatttctg tctagaaatc ctttttgagc attaactgag ttttggtctt  420
cttcctaagt aaccagagaa gatagcagat gtgcacctgt tcacatcttc agtaccacca  480
ggcttcttgg ttgttgatgc rtgcaccagg cttgatttgt caattctcca agtcgtcaca  540
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ggtctgtaaa tgcctactgg gatttttttt tctcttagtt atttctcctt ttttgccttc  660
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gttcaattct gaggttttaa gttgctttta ggctctcact agtctgatct tagctttgta  840
caatgcatgc tcactgtgca tcctgagcct aggctctga tccagagtgg gaacatggag  900
acttctcagc gttcattcct ttaacaggcc ctcacgtttg ctatgtgcag agccttttgg  960
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&lt;210&gt; 205

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens



173

<220>  
 <221> allele  
 <222> 501  
 <223> 12-490-312 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-490-312.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-490-312.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 189..209  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 621..641  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-490-312 potential probe

<220>  
 <221> misc\_feature  
 <222> 437,543,597,604,664,732,872,932  
 <223> n=a, g, c or t

<400> 205  
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 ccgcgacagg ccggctctcc gagcatgtc agcttggect cacgactgca cggcgtcccc 180  
 ccatgcggat ctccacggtc caggttttcc ggacggctct gggggcgggg cctgtgttgg 240  
 tccgcggggc tgcgaggcga ggttgcggtg ccagggcgcc gctgggcaga gatgggttcc 300  
 tggagagcgg gtctcgcgtt tttccgcgga agacactact ggggaatcggc ctttggtcat 360  
 ggtctaggga agagctgatt ctgcaagata tgagctcctt ctttccccctt cagctatcgt 420  
 tacagccctg atggagnctg tctgctgaaa actcacttcc agatgtcgaa acctggccct 480  
 ggggtgcagtc tacagaatga watcgagcag tagtggacct atgcaggcaa agcattgtca 540  
 ganggggagt aactcttgtt aaatatgact tcagttgatc cgtgatggga tttttgnttt 600  
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 ctgnccctgt ggtgcagcag cgccggcgcc ctttttgga ttgtccaaag ctctgggacc 720  
 cccgctgggg gnagctggtt cagaattcgg gggggtagga tagggtagg gaggtccctt 780  
 atgtgagtgt gagccgggat ctaggacaga gtaactccaa ttacagtgtt aaactcttac 840  
 taggttcagt gtcgcgtagg gctttggcta gnactagaat ttagtctatt attgaaccgc 900  
 ccaggtccat gattcctaac aggtgcatgg gnaaaagtgt atggtgtgca gcagggtgat 960  
 gggaaaactg ctttgcaaag tgtaaatcaa ggtctacatc a 1001

<210> 206  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-491-295 : polymorphic base G or A

```

<220>
<221> misc_binding
<222> 481..500
<223> 12-491-295.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-491-295.mis2, potential complement

<220>
<221> primer_bind
<222> 777..795
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 266..286
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-491-295 potential probe

<220>
<221> misc_feature
<222> 128,300,307,312,357
<223> n=a, g, c or t

<400> 206
tgcaggcaga aatgaaaaaa gaaatcctga aactttaaga atcggggata atatcagata      60
agtatgttgc ttaaatttga ctaattttaa ccatttttat aaattagtca ctcagcatga      120
agacatcnaa aaattcttcc taaattctaa acactatgta catttgaatg ctgagttagt      180
cccaggattc atccccaaaa tatcactgac cagtatgatt taatattagt atttgccctc      240
tcaggggcat ccttgctgca ggatatcaca agggtagaag gtaaagcagc tgcacaaaaan      300
catcttntgc tncctgtttg ttactctgag atgacccctc gcctgaaaat atctaangca      360
gatacttttc tcagctttgc aatgcaaatg atatttaaca gtccctaggg aatgttttgt      420
gtgcttatca aatttcagct aaacacattt atttatatga taaatatgat aacccaaaagg      480
ttccatgtat tatcttttca rtattcatgt tatactgtac aagcactgta ttatacaaat      540
actgtataat atagtattgt aaggtacaaa aatacaataa actaaccata cttcaactct      600
gcgggggttgc tgtgggagag ctctgcaggg ttgctgtgca agccccaggt aagtgattaa      660
gtcagacaat aactacatca gtgcttctca aattttcatg tgccctgcaga ttatcggttg      720
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ctgcgtctct cccaagcttt caggtgatac agatgctgac tgggtggacca cattttgagt      840
agcaagatgc taaagcttct tacaactcta aactttcaaa atcctttata aaaggatagc      900
tataatgtat atgtgaaatg gactatggaa acataaaaata ataaaagctt actattgtca      960
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<210> 207
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-493-417 : polymorphic base G or C

<220>
<221> misc_binding
<222> 481..500

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175

&lt;223&gt; 12-493-417.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-493-417.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 90..109

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 514..534

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-493-417 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 590,704,1000

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 207

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catgaaaata	aaaaagtcg	gttggtcctg	tctggggaga	aagttcttga	aactctgcaa	120
gagaaggggg	aaaggacaaa	cccaaagtaa	gctctcatgt	gctgatttag	tgtttctgaa	180
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aaaaaatgta	ttgcagaagc	stataactgg	gatcacaaac	catgatgcaa	gatataagtt	540
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cactctgtcg	tccgggctgg	agtgcagtgg	cacgatctca	gctcactgca	agctccacct	660
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aagtagacag	agacggggag	gggaggggaa	tatatactcc	tctgctcatt	ttccactcat	960
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&lt;210&gt; 208

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-494-373 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-494-373.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

176

```

<222> 502..521
<223> 12-494-373.mis2, potential complement

<220>
<221> primer_bind
<222> 127..144
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 571..591
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-494-373 potential probe

<220>
<221> misc_feature
<222> 2,4,8,15,19,33,36,296,311,640,899
<223> n=a, g, c or t

<400> 208
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gagtgattca atcactcggt tgccccactt gaaaaaaaaa aaaagctgga gtacatatct      180
agaatcctta ccacacgaac attctatatt tttggttctt tgactctaaa atggtcagct      240
taaaagagtg ttttccatcc ctcttttggc tgttagagcc atccctgggtg actttnaccc      300
ctttcccctg naaccctttca tatgtttatt gagcatccac tccagttgtg cactggtggt      360
tgttttcaag gggttactgg taccctccac cctcaccctt ctttctccct gcagaaagct      420
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tacttaccaa atgcttagaga ytgacacagg ctatgtcatt ttttcaatac caagccctgt      540
gaaatgggta ctatcagacc attttttacac ctgtggctta gaagacttaa atgtctagtg      600
taagggcggt tgtagtggta ggcagaggca ggtctcaaan ccctcgcta cctaactcag      660
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gattgttcca ttttatatat atatgaacat gatagacaga cagatagata gataaataga      780
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gagccttgca attccagcat ctccctcatg gtggcagtag aagactaagt catgtacanc      900
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<210> 209
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-495-166 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-495-166.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-495-166.mis2, potential complement

<220>

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177

<221> primer\_bind  
 <222> 336..355  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 784..802  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-495-166 potential probe

<220>  
 <221> misc\_feature  
 <222> 146,405,641,702,965  
 <223> n=a, g, c or t

<400> 209  
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 gtggtaggca gaggcaggct tcaaanccct cgctaccta actcagacac acttggcaat 180  
 actatatatc tctttaagcc ttccagagga tatgttttag gaggtggatt gttccatttt 240  
 atatatatat gaacatgata gacagacaga tagatagata aatagataga gataatttta 300  
 aacgtgctca ctttcattta tgaaaattta catttgataa ctaaaagagc cttgcaattc 360  
 cagcatctcc ctcatgggtg cagtagaaga taaatcagct tttgggtcaag attggtgtta 420  
 caacaactaa ccagagaaga acctccttaa taaatcagct tttgggtcaag attggtgtta 480  
 cgtaaaatgc aaaatttaga ycctcggaag gggaaaaaaa gagtctcaag tatctagggt 540  
 attgacccca gaaacaaaag aaatttgaca aacttgcaaa acagagacaa caacttccta 600  
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 tctgaagggt ttttaacacag cgataactgg catatagaaa tttcacatat ttttcttgat 840  
 cactatggcc tatcaacccc cgctgatga ttaaatgaag gacgatgtat ccaaggttct 900  
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 ttatnatctg ggtacagtag acaccattgg gcagacttca g 1001

<210> 210  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-495-272 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-495-272.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-495-272.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 230..249  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 678..696  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-495-272 potential probe

<220>  
 <221> misc\_feature  
 <222> 40,299,535,596,859  
 <223> n=a, g, c or t

<400> 210  
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 gattgttcca ttttatatat atatgaacat gatagacaga cagatagata gataaataga 180  
 tagagataat ttttaacgtg ctccctttca tttatgaaaa ttacatttg ataactaaaa 240  
 gagccttgca attccagcat ctccctcatg gtggcagtag aagactaagt catgtacanc 300  
 gcagcctcca gcaccaacaa ctaaccaga agaaacctcc ttaataaatc agcttttggg 360  
 caagattggt gttacgtaaa atgcaaaatt aggaccctcg gaaggggaaa aaaagagtct 420  
 caagtatcta ggttattgac ccagaaaaca aaagaaattt gacaaaactg caaaacagag 480  
 acaacaactt cctatggggg waaaaaaaaac aaagcaacta cttgttgctg cccancccg 540  
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 ttttccatt tcctcgggtc ccttgaaacc aagcttttga gaacaaatag ttcatttctc 660  
 ctttcttaat ttgctctgaa ggtttttaac acagcgataa ctggcatata gaaatttcac 720  
 atatttttct tgatcactat ggcctatcaa cccccgcctg atgattaaat gaaggacgat 780  
 gtatccaagg ttcttaaaga acttggttgc aacgaatgaa cacaaaaaac accagacaat 840  
 aaacatgtat tgaattatna tctgggtaca gtagacacca ttgggcagac ttcagaatta 900  
 gcccccggaa gttttgcttt tgcttgactc ttggtcatag ctgattggac tagaagtgaa 960  
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<210> 211  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-495-424 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-495-424.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-495-424.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 78..97  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 526..544  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-495-424 potential probe

<220>  
 <221> misc\_feature  
 <222> 147,383,444,707  
 <223> n=a, g, c or t

<400> 211  
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 ggcagtagaa gactaagtca tgtacanccg agcctccagc accaacaact aaccagaag 180  
 aaacctcctt aataaatcag cttttggtca agattggtgt tacgtaaaat gcaaaattag 240  
 gaccctcggg aggggaaaaa aagagtctca agtatctagg ttattgacct cagaaacaaa 300  
 agaaatttga caaacttgca aaacagagac aacaacttcc tatgggggta aaaaaacaa 360  
 agcaactact tggtgctgcc cancccgttt agtaactggc cttggtagtc ttccagcagc 420  
 aagcacata gccttggtcca cagnatgctt ttcccatttc ctccgtcccc ttgaaaccaa 480  
 gcttttgaga acaaatagtt yatttctcct ttcttaattt gctctgaagg tttttaacac 540  
 agcgataact ggcatataga aatttcacat atttttcttg atcactatgg cctatcaacc 600  
 cccgcctgat gattaaatga aggacgatgt atccaagggt cttaaagaac ttggttgcaa 660  
 cgaatgaaca caaaaaacac cagacaataa acatgtattg aattatnatc tgggtacagt 720  
 agacaccatt gggcagactt cagaattagc ccccggaagt tttgcttttg cttgactctt 780  
 ggtcatagct gattggacta gaagtgaaca cctgactcag gaggaggaag ttcattggct 840  
 gaccaggagc caattagggg ctctctcttg agaactctga ctaagagaca gagatgcagg 900  
 ttatttagtc cagttgcggg tgtagcatta gacagatata aaaccaaga actcttggtgta 960  
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<210> 212  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-500-220 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-500-220.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-500-220.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 283..303  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 711..731  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513

180

&lt;223&gt; 12-500-220 potential probe

&lt;400&gt; 212

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ctgaaatgat	cgctctcctg	ctgcctcagc	cttcaggta	gctggaatta	cagacacaaa	180
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gtgagtattc	tcaccagtag	agcagatggt	gaatatccaa	ggtagtgatt	taaaagacca	360
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taacaaaact	gaatggatca	agttagggtc	aaaagtatgc	cagaagtctt	atctagtgtc	840
ttggtttgtt	tgggctgctg	taacaaaaat	accttacagt	gggtggctta	taaacaacag	900
gaatgtattg	ctcacagttc	tggaggctgg	gaagttcacg	atcaagtcac	cggcagtttc	960
ggtgtgtggt	gagagctttt	gctctctggt	tcatacatgg	t		1001

&lt;210&gt; 213

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-501-155 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-501-155.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-501-155.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 636..655

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 168..188

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-501-155 potential probe

&lt;400&gt; 213

acagctctac	tgagctttac	catctgaaca	cagctaagaa	tttgggattt	taggaacaga	60
actgtaaatc	ttaaaacaag	caaaaagtgc	tttctccttg	gtagaaatac	ctgtggccaa	120
gcccagacct	tcagatccta	acttgggata	caactgattg	ttacttaacc	tgctgctaac	180
ctaaagacta	gttttctgct	catctagtcc	cctgtgtgaa	taagaggttt	cctcctggtc	240
ccaggatgcc	tctgctgctc	agctcatctc	tgaggctgtc	tgtgagcctt	cagttcagaa	300
attgagggaa	aatccattgc	cttggatgtc	tccttccaaa	tcaccaccca	aaacaaaaat	360



181

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taggagacgt agaaatccag gatgtgttgt ggggttaggcc atattctgag gagttggaag 420
gtttcttggg gacctgaagt ctctgaaga gaaaatgtgg aaataagtgt ccttcaagcc 480
attgtagtgc ttgacatggg kgaaagaggc cagggggagg gggacctgtg gtgccagggc 540
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gcatgatatt cagaggccct gctaatagagc tttttgtttg ggtaaagcct cccctgcttt 660
ccctgaagat aagccactga gttctatcag ctgctgtctt ctctccatta tgctttcgta 720
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&lt;210&gt; 214

&lt;211&gt; 998

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-503-52 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-503-52.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-503-52.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 535..552

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 80..100

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-503-52 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 105,135,165,402,424,522,569,587,843,873

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 214

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182

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cattggccat	gctgggtctc	aactcctgac	gtcaagtgac	ccaccgcct	ccgactccca	840
aangtgctgg	gattacaggc	atgagccact	gcngccagct	tgctcacata	cttatttcaa	900
gcttttgaaa	tgtctaacat	ttccctttct	tcacttttgt	cttcacaatt	caaaacagat	960
tctttagtta	actctgaatt	tggttaattgt	tctattct			998

&lt;210&gt; 215

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-503-62 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-503-62.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-503-62.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 545..562

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 90..110

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-503-62 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 115,145,175,412,434,532,579,597,853,883

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 215

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aacttttttc	tgttgatata	wcatcattct	taatgtgaac	atacttaata	antgttacat	540
acttggtcac	aatcgggtac	atctgggtata	taataaaaana	tcaagggtatt	ttgcttnccct	600
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183

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caaaacagat tctttagtta actctgaatt tggtaattgt t                               1001

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&lt;210&gt; 216

&lt;211&gt; 1013

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-504-54 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-504-54.misl, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-504-54.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 446..463

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 993..1013

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-504-54 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 108,180..181,663,712,768

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 216

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184

<210> 217  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-504-96 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-504-96.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-504-96.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 406..423  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 953..973  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-504-96 potential probe

<220>  
 <221> misc\_feature  
 <222> 68,140..141,623,672,728  
 <223> n=a, g, c or t

<400> 217  
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 gcccagcgtg gttgtgctta ttttaaattc ttaatttggt tgttccagta tttctgcttg 300  
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<210> 218  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-504-428 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-504-428.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-504-428.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 75..92  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 622..642  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-504-428 potential probe

<220>  
 <221> misc\_feature  
 <222> 292,341,397  
 <223> n=a, g, c or t

<400> 218  
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 ctgttgcccc tccactaggc attactctgc tagggattct cctctaaagc ctttgaaagg 240  
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 aataaaaagga ggaaggaaac cctgttacta gtttaagtaa gcaataatgg agatctggat 720  
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 gtttattctt agccttttct tttcttttct ttcttttttt ttttttttgc agtgaattct 900  
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<210> 219  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 474

186

&lt;223&gt; 12-507-53 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 454..473

&lt;223&gt; 12-507-53.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 475..493

&lt;223&gt; 12-507-53.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 422..441

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 982..1001

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 462..486

&lt;223&gt; 12-507-53 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 153,560,925,934

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 219

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gagtttctga ttcagtaaat ctggggcaag ccnttgataa tgtgtttcct aacaagtcct      180
caactcatag tgacactact agtccaggga tcaactctga gaatcactgc catataccaa      240
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cagtgtcag gtctgttccc actgattctg atttaattgg ttgggaatga ggccaggata      360
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&lt;210&gt; 220

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-507-92 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

187

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<222> 481..500
<223> 12-507-92.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-507-92.mis2, potential complement

<220>
<221> primer_bind
<222> 410..429
<223> upstream amplification primer

<220>
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<222> 970..990
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-507-92 potential probe

<220>
<221> misc_feature
<222> 141,548,913,922
<223> n=a, g, c or t

<400> 220
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cagtaaatct ggggcaagcc nttgataatg tgtttcctaa caagtcccca actcatagtg      180
acactactag tccagggatc aactctgaga atcactgcca tataccaata tttctcagta      240
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ctgttcccac tgattctgat ttaattgggt gggaatgagg ccaggatacc tgtattttat      360
aaaagttctt caggagattc taagggtatac ttaggttcaa gaagcacttc cttacactca      420
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tcaatggtta gctattatca rgatattttt atatagactg ggatgcctat tagtcttagt      540
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<210> 221
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 500
<223> 12-507-159 : polymorphic base G or T

<220>
<221> misc_binding
<222> 480..499
<223> 12-507-159.mis1, potential

<220>

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<221> misc\_binding  
 <222> 501..520  
 <223> 12-507-159.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 341..360  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 901..921  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 488..512  
 <223> 12-507-159 potential probe

<220>  
 <221> misc\_feature  
 <222> 72,479,844,853  
 <223> n=a, g, c or t

<400> 221  
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 cctggggatt ttaaagagaa tacaactgcc aaaaattacc agtgctcagg tctgttccca 240  
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<210> 222  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-507-177 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-507-177.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-507-177.mis2, potential complement



189

<220>  
 <221> primer\_bind  
 <222> 324..343  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 884..904  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-507-177 potential probe

<220>  
 <221> misc\_feature  
 <222> 55,462,827,836  
 <223> n=a, g, c or t

<400> 222  
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 gaatacaact gccaaaaatt accagtgtctc aggtctgttc ccactgattc tgatttaatt 240  
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 atacttaggt tcaagaagca ctcccttaca ctcaatctcc acccgtcatt cagagaaact 360  
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 tgctcctact ttgccttcca ccatgagtaa aagttccttg aggcctcccc agaagcagac 600  
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<210> 223  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 12-508-29 : polymorphic base A or G

<220>  
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 <222> 481..500  
 <223> 12-508-29.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-508-29.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 473..491  
 <223> upstream amplification primer

<220>  
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 <222> 907..925  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-508-29 potential probe

<220>  
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 <222> 847  
 <223> n=a, g, c or t

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<210> 224  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
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 <223> 12-509-42 : polymorphic base A or G

<220>  
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 <222> 481..500  
 <223> 12-509-42.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-509-42.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 460..479  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 889..909

191

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-509-42 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 678,728,796,824

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 224

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&lt;210&gt; 225

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-509-126 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-509-126.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-509-126.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 376..395

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 805..825

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

192

<222> 489..513  
 <223> 12-509-126 potential probe

<220>  
 <221> misc\_feature  
 <222> 594,644,712,740  
 <223> n=a, g, c or t

<400> 225  
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 ggggtgttta acaaaaagct aaaagtttga agaacctctt tgaggactag tgcagaaaga 180  
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 ctgagggatg ggccagttca tctgttaaaa tctatgactg taactattgt tacagcccag 900  
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 gtttcatcca gtattataac ttttcctcgc ttcatatagg t 1001

<210> 226  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-510-59 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-510-59.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-510-59.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 539..559  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 107..127  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-510-59 potential probe

<220>

193

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<221> misc_feature
<222> 7..8,169,196,281,468,470,821
<223> n=a, g, c or t

<400> 226
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taaactttca tatcancaaa atgtataaat cttaaataata aactcaatga atcacttgag      240
tgagcttaag tgtaagctca aacaataact taatagtttt ncataagtgc tagaaatctt      300
atccaggaat aattgaaatt ttaccaatga gcataatata ccaaataaaa taaatgggct      360
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gcaagggtcat cacaccatgg cctgtgggac taggggctaa taatggtnntn ttatattttt      480
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aaccctggag ctacagagta aaagtcaact ggaagccaag acagtgattht cttttctttg      660
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<210> 227
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-511-74 : polymorphic base T or C

<220>
<221> misc_binding
<222> 502..521
<223> 12-511-74.mis1, potential complement

<220>
<221> misc_binding
<222> 483..500
<223> 12-511-74.mis2

<220>
<221> primer_bind
<222> 558..575
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 125..145
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-511-74 potential probe

<220>
<221> misc_feature
<222> 102,561,637
<223> n=a, g, c or t

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194

&lt;400&gt; 227

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&lt;210&gt; 228

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-325-311 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-325-311.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-325-311.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 191..208

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 596..613

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-325-311 potential probe

&lt;400&gt; 228

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195

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gcaaaggagc agcgccctcac rgacaggctg tgactccctg gcacgggggg acacggagcc 540
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ctctgaggtg atccaggag agggaccagg ctggacaaag gcctggaggg ttcagtggag 660
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&lt;210&gt; 229

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-327-120 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-327-120.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-327-120.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 382..400

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 805..824

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-327-120 potential probe

&lt;400&gt; 229

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<210> 230  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-331-179 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-331-179.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-331-179.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 326..345  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 728..747  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-331-179 potential probe

<400> 230  
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<210> 231  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501



197

&lt;223&gt; 10-331-357 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-331-357.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-331-357.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 148..167

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 550..569

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-331-357 potential probe

&lt;400&gt; 231

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cacaggtttt ggggaattagg atgtgggtgg atctttctgt ggtgggggggt gggggcaaca	180
ttcaacccat tacatagggg gacccaacc aacctgtgcc ccaacctctc tccagggttc	240
aacttctcaa cagagtctat ggccaggcct gaaggagggt tgaacgtgta ccaccacctt	300
gtagagacgc tcaagtttgc caaggggcag aggtggaggc tgggggaccc tcgaagccac	360
ccgaagctcc aggtgaggtt gctgaggttg ctgggctggt gggccgtcct cctccctggc	420
tcaggacttg gcatgaaatg agggtcaggc ctggtagggg gaagtggag ggatatgtat	480
gtggttctag gccagggcag kactgaaagg gatcccgggg tggcaggtag aggggtcagg	540
tgcaggagtg gcaccatata tcaaaggacc tggagggtga gcagagtcta gacctagctg	600
ggcttgaggg agacctggcc acaaggtaga ggacagactg gaggtggccc ccatgggggc	660
tgatctcatc ctgcccttgg ttctgctggg tctgcctggc ccctcactga ccctgccacc	720
tgccacccca cccagaaatg cctcccgga cctgctgggg gagaccctgg cccagctcat	780
ccgccaacag atcgatggcc ggggggacca ccagctcagc cactacagct tggccgaggc	840
ctggggccac gggacaggca cgtcccatgt gtctgtgctg ggggaggatg gcagcgccgt	900
ggctgccacc agcaccatca acacaccgtg cgtagggcct gggggaaggc ggatggcttc	960
actcctctc tcctagacct gcacaccccc agccccatgt c	1001

&lt;210&gt; 232

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-334-263 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-334-263.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

198

<222> 502..521  
 <223> 10-334-263.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 240..257  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 658..675  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-334-263 potential probe

<400> 232  
 tgcacacccc cagcccatg tcccctcact tgtccccacg gggcagcacc ttgcttttgc 60  
 cctttttctc ctctctatt tcaaaagagg cccccacccc tgacatctct ggctggaaag 120  
 gctgctgctg ggggtggccc gacccaagat ttacctggga atgggtagcc tctactcagaa 180  
 ggggtgccctg atgtgggggc acaggtgggt ctttggggac ccctcctggg tgggtgccagg 240  
 gagagaatag cggcttcagc atgcttcggg gcagctgtaa aacgaggggg tcctgcaaag 300  
 cgtgcagggt aagtgggtgc tgggtgggag cccgggtcct agcccaggct cttctgcctc 360  
 cacggctgca gctttggagc gatgggtgtat tcaccacgga caggcatcat cctcaacaac 420  
 gagctcctgg acttatgcga gcgatgcccc cgggggttccg gcaccacccc ctacacctgg 480  
 gagaacaaaag cttcccaccc rgggtccaca agggccccc accaggggag aggagggagg 540  
 gggctgggct ggggttgcat gctaaccctt ggatgggtca ctgcacttgc caagacgctg 600  
 tttgctcagc agtgagtggg gacaggggtg gtggagctcc cggaagggtg tggccccag 660  
 ttccaggcga gcgttcccca tctccatgg tgcctccat cttgatcaac aaagcccagg 720  
 ggtcgaaagt agtgattggc ggggctggcg gggagctcat catctctgct gtggcccagg 780  
 tgagtctggg gctcctggct cgagtgtctc ctctctgggc agcatactgt ctgactgtct 840  
 ctggagtggg gatgtgagg ctgatgtagg gtagcagggt gcccccttct tccctgaaac 900  
 cctcatctct ccccaggcc atcatgagca agctgtggct tggctttgac ctgagagcgg 960  
 ccattgcagc ccccatcctg catgtcaaca gcaagggtctg t 1001

<210> 233  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-321-226 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-321-226.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-321-226.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 276..293  
 <223> upstream amplification primer

<220>

199

<221> primer\_bind  
 <222> 692..711  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-321-226 potential probe

<400> 233  
 gcacaggtgg gtctttgggg acccctcctg ggtggtgcca gggagagaat agcggcttca 60  
 gcatgcttcg gggcagctgt aaaacgaggg ggtcctgcaa agcgtgcagg gtaagtgggtg 120  
 tctggtggga gccccgggtc ctagcccagg ctcttctgcc tccacggctg cagctttgga 180  
 gcgatggtgt attcaccacg gacaggcatc atcctcaaca acgagtcctt ggacttatgc 240  
 gaggatgcc cccgggggtc cggcaccacc ccctcacctg gtgagaacaa agcttcccac 300  
 ccagggtcca caagggtccc ccaccagggg agaggaggga gggggctggg ctgggggttg 360  
 atgctaaccc ctggatgggt cactgcactt gccaaagacg tgtttgctca gcagtgtgtg 420  
 gagacagggg ggggtggagct cccggaaggt gctggcccc agttccaggc gagcgttccc 480  
 catcctccat ggtgccctcc rtcttgatca acaaagccca ggggtcgaag ctagtgtattg 540  
 gcggggctgg cggggagctc atcatctctg ctgtggccca ggtgagtctg gggctcctgg 600  
 ctgagtgtc tctctctgg gcagcatact gtctgactgt ctctggagtg gggatgtgag 660  
 ggctgatgta gggtagcagg gtgccccctt tctccctgaa accctcatct ctccccagg 720  
 ccatcatgag caagctgtgg cttggctttg acctgagagc ggccattgca gccccatcc 780  
 tgcatgtcaa cagcaagggc tgtgtggagt acgagcccaa cttcagccag gtgaggctga 840  
 ggtccgagct ggatgcctag ggcagagccc actccccaaa tccgtgtctg tcaaagccac 900  
 ctgggaggaa ctcagtcact gagattctta ggccagggtac acttcaactt tggggggccat 960  
 aggagttggg gaccttgatg ggtgaggctg tcagtggcct c 1001

<210> 234  
 <211> 858  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-183-98 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-183-98.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-183-98.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 581..598  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 136..155  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-183-98 potential probe

200

```

<400> 234
ctgagacccc agaggggtccc tcccagcatc ttcaaagcaa caggattttg tgcctgcaga      60
tcctttctttg cagcacacac caccaccctt gaccaggacc cctagaatgc ccagcatccc      120
tgaggaggcc ctgtggtagt ttcagctccc tctgggggcc cagaatgaac ctggcctgtg      180
gtgaggatgt aagcaccaat ggccaattgg gtccaaagga agacaccggt tcaaacactg      240
aaaccaatca gattctccca cggecttctt gccatcagac gacactgggt caggggtggt      300
tgctatgtac agggcagagc cacccaatcc ccacgcagga gctgtgtcct gccatgctgg      360
cctctctctg gccatcacat caggccaagc aggggagagg aatgggaatg cccatgcacc      420
cctatcaact ctgcagacac agaaccatgc acagctcttg ggaggagtca gatgagctgc      480
tcaaagcccg ggagggaccc rcacagtggg cagcatagca gggacggtgc tttagccaag      540
gcagggatgg caggtgactc actcaggatc ttcaaaggag ccgctgcatt tccgtgctct      600
ttccagataa gaaggacatg tcggtgatga tgagcgagac ggacatgaac gccatcgag      660
gcacgctgaa gctgtacttc cgtgagctgc ctgagccccct cttactgac gaggttctacc      720
ccagcttcgc agagggcatc ggtgagcatt ggaggccttg gcctcgtggg agacgtctcc      780
tccatgtgca ctgctgccct tggaggctgt gaaaagtga ggtgtgggaac ctgagctgtg      840
accctctgc catgggtcg                                     858

<210> 235
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-185-78 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-185-78.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-185-78.mis2, potential complement

<220>
<221> primer_bind
<222> 424..444
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 855..875
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-185-78 potential probe

<220>
<221> misc_feature
<222> 661,672,954
<223> n=a, g, c or t

<400> 235
gctgagaaag ccacagttct ggcagctggc agagctgaaa gagaagggaa aatctatgca      60
cagggccctt ggatttccac ccatattccc caggaaagga tacagttcca ggaagggctc      120
tctctgatct cccggaatga aatctgagat gaaccacagg gccttatgtc ttacctgtac      180
ctcctttttg tagtttccag ctggcctggc acagactgac aaccccaaat ggatggctga      240
aatccactgt tctccagtcc ctctcaaacc tcaccggcaa gaaggaaaga aactaaaccc      300

```

201

cgagggccca	ttatattcca	gcggggctct	agggattcca	cactcattca	gttcctttac	360
atcataagaa	gtgggtatta	taatatcaga	tccactgtaa	aaggatgatcc	cgaagcaaat	420
ggtagagcag	gagctgaat	aggggcacct	ccaacagcca	catccacccc	accacagctc	480
ccaggggttt	ctttctaata	ycagggtgg	cctataccct	gggactatag	ggggccttgg	540
tgtgctaaat	cttcagagac	atttttaaac	ggcaagacg	cctcttggcg	actggaagaa	600
gttcctgacc	tcgcggcatt	ccctccttcc	catccccact	accacaccaa	tcctatgcca	660
ngcctgtctg	tngtcaggga	agaagtcag	ctggacaact	ctggcagcct	cctgtcgccc	720
tcccctgcca	gcaatccttc	tgtgccacag	ctgtgtgtcc	acactctaaa	gcacacatcc	780
caccaggcca	ctctcctgta	aaataatgcc	tactgtccc	aggccaaata	ccccagcgcc	840
gatacctgct	tcttcaaata	caggctccaa	cccacccctc	catctcattt	ccatttttacg	900
tgcacacata	gtcacacact	ctcacatacc	cacactttgg	ctcctactgt	tcctatcccc	960
tgaaaacccc	ttaccacttt	cctccctgag	aaataactgc	t		1001

&lt;210&gt; 236

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-186-154 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-186-154.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-186-154.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 348..368

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 784..803

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-186-154 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 3

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 236

ttntttttct	tttttttgag	acggagtctc	gctctgttgc	ccaggctggc	gtgcagtggc	60
acgatcttgg	ctcactgcaa	gctctgcctc	ccgagttcac	gccattcccc	tgccctcatcc	120
tcccgccctca	tcctcccaag	tagctgggac	tacaggcacc	caccaccacg	ctcggctaata	180
tttttgtatt	tttagtagag	acagggtttc	accgtgttag	ccaggatggg	ctcgatctcc	240
tgaccttgtg	atccgcccgc	cttggcctcc	caaagtgtctg	ggattatagg	cgtgagccac	300
cgcgcccggc	ccctcccttg	actcttgact	gaaggacctt	tgtctttgtg	aacatcaatt	360
ctcaggacct	ttcacccctg	ggacgtgaaa	tgctgagaat	ttgggagatg	acagtctggg	420
gactgggatt	aatggaatcc	agtgaccac	aaacctaagg	ttctcagctc	ccttggggag	480
ttggaatgtc	agctattcag	rtctagggct	ttccatggag	taaatcctaa	actctggggt	540

202

```

ggagacttta agcctccaag gaccttcaca gctaaggccc agggactagg gcgaggagag 600
tctttgatcc tcagagtctt ggagtttggc cagtggactc tgaggaatgg agtctctgag 660
cactgaaggt ccaacttttg cttcagcagt aaaggatctt ggccttcaag tctaaggaca 720
gtgggcaatt agtaggtcag gcatggggaa ctcatagcca aacgtgcagg gctccaaaga 780
cctcatttgc cctgtcagca gctcaggcca tgtggcatca cccgatgcat ctagatgtct 840
ccggaatctc aggccttgca ggggtgaggtt ctcgggccat tagttttttt tgttttgttt 900
tgttttgttt ttttggttg ttgttggtga gacaaagttt cactctgtca ccaggtctgg 960
agtgcagtgg cgcgatctca gcttattgca acctccacct c 1001

```

&lt;210&gt; 237

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-186-397 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 12-186-397.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-186-397.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 105..125

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 541..560

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-186-397 potential probe

&lt;400&gt; 237

```

ccttgtgatc cgcccgctt ggcctcccaa agtgctggga ttataggcgt gagccaccgc 60
gcccgggccc tcccttgact cttgactgaa ggacctttgt ctttgtgaac atcaattctc 120
aggacctttc accctgggga cgtgaaatgc tgagaatttg ggagatgaca gtctggggac 180
tgggattaat ggaatccagt gaccacaaa cctaagggtc tcagctccct tggggagttg 240
gaatgtcagc tattcaggtc tagggctttc catggagtaa atcctaaact ctgggttgga 300
gactttaagc ctccaaggac cttcacagct aaggcccagg gactagggcg aggagagtct 360
ttgatcctca gagtcttgga gtttgccag tggaactctga ggaatggagt ctctgagcac 420
tgaagggtcca actttggctt cagcagtaaa ggatcttggc cttcaagtct aaggacagtg 480
ggcaattagt aggtcaggca yggggaactc atagccaaac gtgcagggct ccaaagacct 540
catttgccct gtcagcagct caggccatgt ggcatacccc gatgcatcta gatgtctccg 600
gaatctcagg ccttcgaggg tgaggttctc gggccattag tttttttgt tttgtttgt 660
tttgtttttt tggtctgttg ttgttgagac aaagtttcac tctgtcacc aggctggagt 720
gcagtggcgc gatctcagct tattgcaacc tccacctcct gggttcaagc aattctcatg 780
tctcagctc ccaagtagct gggattacaa gtgtgtgcca ccaagcctgg ctaatttttg 840
tatttttagc agaaacagcg tttctccatg ttggccaggc tggtctcaaa ctccctgacct 900
cagggtgatc gccaccttg gcctcccaaa gtgctgggat tacaggcatg accaccgcgc 960
ctggctaggg agcagtgttt ttaaggacaa cttggcggtt t 1001

```

203

<210> 238  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-187-65 : polymorphic base C or T

<220>  
<221> misc\_binding  
<222> 482..500  
<223> 12-187-65.mis1

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-187-65.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 437..456  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 839..859  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-187-65 potential probe

<400> 238  
cctatatatttt tattttattta tttattttatt tttctgagat ggagtctcac tctgtcacct 60  
aggctggagt gcagtgggtgc aatcttggct cactgcaacc tccgcctccc aggttcaagg 120  
gattcttgtg cctcagcctc ccgagtaaat gggattacaa gcacccgcca ctgcatctgg 180  
ctaattttttg tatttttagt agagacaagg tttcaccacg ttggccaggc tgggtctcgaa 240  
ctcctaacct caaatgatct gccggcctca gcttcccaaa gtgctgggat tacagggtgtg 300  
agccaccgtg cccggccggc atcctatatt tttatatctt aactctggca accttaaata 360  
taagtgacaa atgaaaccaa aaataatggg caaaggtcta aacacatcag tgatcataaa 420  
aaggaattga gaacattgtc agaacactgt cctgccttaa acactcttct ggaaagtcc 480  
acaggttaag tgtgtcgtgt ygtcaggacc tgagatctct tctgccaatt aagctcccac 540  
accatagaat aaggaggaaa atttgcacat ctgctttatg aagccagcgt agcactgacc 600  
ccaaacttgg caaagaaaaac cgcagactaa ttgtgtttat gatgcaaaagc aaaatcctaa 660  
caaaattgga gaatatataa ttaaaaaaga gaatccagta gtacattaaa aagaacaaca 720  
tgacttacct accctagtgg agccaacgct aatattgcaa ggaagatctt ctgagtatgt 780  
ccctgaccgt gactcttagc tcttaccttt caccagagga gaaaaaaatc actccatcga 840  
gactgaagac gcatttgaca atatttaaca cccattcttt tttttttttt ttttttaaaga 900  
caggctcac tctgttggc atgctggagt ggagtgggtgc gatcacagct cactgcagcc 960  
ttgacctct aggtcaaga gatcctcctg cctcagcttt t 1001

<210> 239  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-187-66 : polymorphic base A or G

```

<220>
<221> misc_binding
<222> 481..500
<223> 12-187-66.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-187-66.mis2, potential complement

<220>
<221> primer_bind
<222> 436..455
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 838..858
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-187-66 potential probe

```

```

<400> 239
ctatatTTTT atttatttat ttatttattt ttctgagatg gagtctcact ctgtcaccta    60
ggctggagtg cagtgggtgca atcttggtct actgcaacct ccgcctccca gggtcaaggg    120
attcttgtgc ctcagcctcc cgagtaaag ggattacaag caccgcccac tgcattctggc    180
taatttttgt atttttagta gagacaaggt ttcaccacgt tggccaggct ggtctcgaac    240
tcctaacctc aaatgatctg ccggcctcag cttcccaaag tgctgggatt acaggtgtga    300
gccaccgtgc ccggccggca tcctatatatt ttatatattt aaaggtctaa acacatcagt gatcataaaa    360
aagtgcacaaa tgaaacccaaa aataatgggc aaaggtctaa acacatcagt gatcataaaa    420
aggaattgag aacattgtca gaacactgtc ctgccttaaa cactcttctg gaaagttcca    480
caggttaagt gtgtcgtgtt rtcaggacct gagatctctt ctgccaatta agctcccaca    540
ccatagaata aggaggaaaa tttgcacatc tgctttatga agccagcgta gcactgaccc    600
caaacttggc aaagaaaacc gcagactaat tgtgtttatg atgcaaagca aaatcctaac    660
aaaattggag aatatataat taaaaaagag aatccagtag tacattaaaa agaacaacat    720
gacttaacct ccttagtgga gccaacgcta atattgcaag gaagatcttc tcagtatgtc    780
cctgaccgtg actcttagct cttacctttc accagaggag aaaaaaatca ctccatcgag    840
actgaagacg catttgacaa tatttaacac ccattctttt tttttttttt ttttaaagac    900
aggtctcact ctgttgccca tgctggagtg gagtgggtgcg atcacagctc actgcagcct    960
tgacctecta ggctcaagag atcctcctgc ctcagctttt g                                1001

```

```

<210> 240
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-189-348 : polymorphic base G or A

```

```

<220>
<221> misc_binding
<222> 502..520
<223> 12-189-348.mis1, complement

```

```

<220>
<221> misc_binding
<222> 481..500

```



205

<223> 12-189-348.mis2, potential

<220>  
<221> primer\_bind  
<222> 832..849  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 384..402  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-189-348 potential probe

<220>  
<221> misc\_feature  
<222> 69,615  
<223> n=a, g, c or t

<400> 240  
cgaggcaggt ggatcaagtg aggtcagaag tttgagacca gcctagccaa catggtgaaa 60  
ccctgtctnc taccaaaaaat ataaaaaatt agctgggtgt ggtggtgcgt gcctgtaatc 120  
ccagctaatt gggaggctga ggcagaagaa tcgcttgaac ccgggaggca gaggttgcaa 180  
tgagccgata tcgtgccgct gcactccagc ctgggcaaca aagtgagact ccctctcaaa 240  
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caaatgagg acttgtcggc aagggccaga agaccagtca agccctgggc ttcattgtgac 600  
acaggacggt gaatnaaaaag ccgtgcagtc ctataacctg acttgcagtt cccaaagact 660  
aacctgcgct gctgtactga agatacagaa tgggagggtc agggatgagg aaaacagaat 720  
gggcatctaa ttccctgtaa tgtctttgct ctcccttaat ctaagcttta tgttcaaaat 780  
aaaagtacca ctctcactcc acaactcccc cagtatctga agaccaccat ggcgattccc 840  
taaattggcc ctcttccaa gtcagcacca cacattcccc atccactcct ccagatttca 900  
gtaccctgtg tccactggca tcaaagtatg acaggcacct cggaagtggg cacagatata 960  
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<210> 241  
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<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-192-63 : polymorphic base C or G

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-192-63.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-192-63.mis2, potential complement

<220>  
<221> primer\_bind

206

&lt;222&gt; 544..563

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 75..94

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-192-63 potential probe

&lt;400&gt; 241

caagtttccc	ctgctgccac	gcaggcgagg	ggagcctttg	atgctgccac	ctcccttaga	60
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catcaacagt	gcactgcagg	ttgaggacaa	ggccatctcg	gactgcagac	cctcacggcc	180
ttcccacact	ttgtcctcac	ttgcaacagg	ggcttctggt	ctgcctgccg	tttctaaagc	240
accagtatg	gatgcacagc	aggagacaca	caagtcccaa	gactgcctgg	gcctactgga	300
ccccttagca	tctgctgcag	gggtccccctc	tacagctccc	atgtctggga	agaagcacag	360
accaccaggc	cccctgttct	cctcctcaga	tccccttctt	gccacctctt	ctgattccca	420
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ttctgggcct	cagccacagc	tgcagcaggt	gcccagaggt	cagaaccaga	gatcccagac	660
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aggcctcccc	gaacaaaagc	ggaagagggg	ccagcctcat	cccactgcca	gctgaccctc	780
agttcctcaa	acacagttag	tgaggacgga	cctcaggctg	tctcttcggg	tcacacccag	840
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agatcccagg	cctctaggcc	ccgtatatgc	aagtttcccc	tgctgccacg	caggcgaggg	960
gagcctttga	tgctgccacc	tccttagag	atgggggtacc	g		1001

&lt;210&gt; 242

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 502

&lt;223&gt; 12-192-64 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..501

&lt;223&gt; 12-192-64.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 503..522

&lt;223&gt; 12-192-64.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 545..564

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 76..95

&lt;223&gt; downstream amplification primer

&lt;220&gt;

207

&lt;221&gt; misc\_binding

&lt;222&gt; 490..514

&lt;223&gt; 12-192-64 potential probe

&lt;400&gt; 242

```

acaagtttcc cctgctgcca cgcaggcgag gggagccttt gatgctgcca cctcccttag      60
agctggggta ccgggtcact gttgaagacc tggaccggga gaaggaggcg gccttccagc      120
gcatcaacag tgcactgcag gttgaggaca aggccatctc ggactgcaga ccctcacggc      180
cttcccacac tttgtcctca cttgcaacag gggcttctgg tctgcctgcc gtttctaaag      240
caccacagat ggatgcacag caggagacac acaagtccca agactgcctg ggcctactgg      300
accccttagc atctgctgca ggggtcccct ctacagctcc catgtctggg aagaagcaca      360
gaccaccagg cccctgttcc tctcctcag atccccttcc tgccacctct tctgattccc      420
aggactcagc ccaggtcacc tcgctgattc ctgccccctt cccagctgca agcatggatg      480
cgggcatgag aagaacaagg cytggcactt ctgctcctgc agctgccgca gcagccccctc      540
cccgctccac attgaacccc acgttggggg cactactgga gtggatggag gcccttcaca      600
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cctcctggac cagctcgtgc cccaaatgaa atgccatctc gagccccctac agctctacgg      720
gaggcctccc ggaacaaaag cgggaagagg gccagcctca tcccactgcc agctgaccct      780
cagttcctca aacacagtga gtgaggacgg acctcagggt gtctcttcgg gtcacaccca      840
gtgtgaaaag acggcagata cagcaccagg gcagacactc gcctccaggg gtgggtcccc      900
cagatccccc gcctctaggc cccgtatatg caagtttccc ctgctgccac gcaggcgagg      960
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```

&lt;210&gt; 243

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-192-268 : polymorphic base C or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-192-268.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-192-268.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 749..768

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 280..299

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-192-268 potential probe

&lt;400&gt; 243

```

tgccggggct aaagcgagg agggggccag cctcatccca ctgccagctg accctcagtt      60
cctcaaagac agtgagttag gacaggcctc aggctgtctc ttcagggtcac acccagtgtg      120
aaaaggcagc agatatagca ccagggcaga cactcaccct caggaatgac tcctccacat      180
ccgaggcctc tagggccagt acacacaagt ttcccctgct gccacgcagg cgaggggagc      240

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208

ctttgatgct	gccacctccc	ttagagctgg	ggtaccgggt	cactgttgaa	gacctggacc	300
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tctcggactg	cagaccctca	cggccttccc	acactttgtc	ctcacttgca	acaggggctt	420
ctgggtctgcc	tgccgtttct	aaagcaccca	gtatggatgc	acagcaggag	acacacaagt	480
cccaagactg	cctgggccta	stggaccctt	tagcatctgc	tgcaggggtc	ccctctacag	540
ctcccattgc	tgggaagaag	cacagaccac	caggccccct	gttctcctcc	tcagatcccc	600
ttcctgccac	ctcttctgat	tcccaggact	cagcccaggt	cacctcgctg	attcctgccc	660
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tctcgagccc	ctacagctct	acgggaggcc	tcccggaa	aaagcggaag	agggggccagc	960
ctcatcccac	tgccagctga	ccctcagttc	ctcaaacaca	g		1001

&lt;210&gt; 244

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-192-334 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-192-334.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-192-334.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 815..834

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 346..365

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-192-334 potential probe

&lt;400&gt; 244

cctcctggac	gagctcctgc	accaaccgaa	atgccatctc	cagctcctac	agctccacgg	60
gaggcttgcc	ggggctaaaag	cggaggaggg	ggccagcctc	atcccactgc	cagctgaccc	120
tcagtccctc	aaagacagtg	agtgaggaca	ggcctcaggg	tgtctcttca	ggtcacaccc	180
agtgtgaaaa	ggcagcagat	atagcaccag	ggcagacact	caccctcagg	aatgactcct	240
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acaagtccca	agactgcctg	ggcctactgg	accccttagc	atctgctgca	ggggctccct	600
ctacagctcc	catgtctggg	aagaagcaca	gaccaccagg	ccccctgttc	tcctcctcag	660
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209

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ctgctcctgc agctgccgca gcagccccctc cccgctccac attgaacccc acgttgggggt      840
cactactgga gtggatggag gcccttcaca tttctggggc tcagccacag ctgcagcagg      900
tgcccagagg tcagaaccag agatcccaga cctcctggac cagctcgtgc cccaaatgaa      960
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<210> 245  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-192-352 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-192-352.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-192-352.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 833..852  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 364..383  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-192-352 potential probe

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<400> 245
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cccactgcc a gctgacctc agttcctcaa agacagttag tgaggacagg cctcaggctg      180
tctcttcagg tcacacccag tgtgaaaagg cagcagatat agcaccaggg cagacactca      240
ccctcaggaa tgactcctcc acatccgagg cctctaggcc cagtacacac aagtttcccc      300
tgctgccacg caggcgaggg gagcctttga tgctgccacc tcccttagag ctggggtagc      360
gggtcactgt tgaagacctg gaccgggaga aggaggcggc cttccagcgc atcaacagtg      420
cactgcaggt tgaggacaag gccatctcgg actgcagacc ctcacggcct tcccacactt      480
tgtcctcact tgcaacaggg rcttctggtc tgctgcccgt ttctaaagca cccagtatgg      540
atgcacagca ggagacacac aagtcccaag actgcctggg cctactggac cccttagcat      600
ctgctgcagg ggtcccctct acagctccca tgtctgggaa gaagcacaga ccaccaggcc      660
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gaacaaggca tggcacttct gctcctgcag ctgccgcagc agccccctcc cgctccacat      840
tgaaccccac gttgggggtc ctactggagt ggatggaggc ccttcacatt tctgggcctc      900
agccacagct gcagcaggtg cccagaggtc agaaccagag atcccagacc tcctggacca      960
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<210> 246  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-194-135 : polymorphic base G or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-194-135.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-194-135.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 363..381  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 878..893  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-194-135 potential probe

<220>  
 <221> misc\_feature  
 <222> 223,412,431,439..440,565..566,894  
 <223> n=a, g, c or t

<400> 246  
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 gctcgggagg ccgagacagg agaattgctt aggaggcaga ggggtgcagtg aaccgaggtc 180  
 tcgccactgc actccagcct gggcgacaga gtgagactct gtntctcaaaa aaaaaaaaaa 240  
 acagctatta caatatacat gtatgttgta aaataaccaa accaacatag aagggtatat 300  
 agtgaaagga gaagtattgt tgtgggtaaa attgtttcaa taaatggcta atccagtatt 360  
 tttttcctcc aaaaaaacac cttttcccct ttgatttagt gtataaccaca cnagagtcgc 420  
 ttagccgatt ncagtgacnn ttttttgata gctggagaat gcttcctaata ggttgcagggt 480  
 gtgtagggtt tttcttgaca ktttcagtaa gaaagtgaat gttgtgccat aggaaagctt 540  
 tatcacaggt gcacgttgggt agccnnaact aaataacgtg agtgtcaagt acagtatcta 600  
 gtgagtgagg aataactgta ataattacaa ttacagttgt agttaaagtg aagtttgtat 660  
 caaaatttct gtttcaaagg tgaactttta ggaggtgtat ctgcggtgtt tcttcctcag 720  
 atgtgatttc tctgaagcag tggttctcta taactgactt gacacacagt gcatctgata 780  
 tttcgaatga cttaagtgat agtatcaaaa ttactatcat atacatattt aaatcacatg 840  
 actaacctgt gtgtggtatt atgcaaactg ctcagctcag taagggaaca gaanataaga 900  
 aaaaaaggac aatattggga gtcttttata caattcaatt tattcaattg aaggactatc 960  
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<210> 247  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501

211

&lt;223&gt; 12-194-325 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-194-325.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-194-325.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 171..189

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 686..707

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-194-325 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 31,220,239,247..248,373..374,702

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 247

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tcttgacatt	ttcagtaaga	aagtgaatgt	tgtgccatag	gaaagcttta	tcacaggtgc	360
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tgaagcagtg	gttctctata	actgacttga	cacacagtgc	atctgatatt	tcgaatgact	600
taagtgatag	tatcaaaatt	actatcatat	acatatttaa	atcacatgac	taacctgtgt	660
gtggtattat	gcaaactgct	cagctcagta	agggaaacaga	anataagaaa	aaaaggacaa	720
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taaagagagc	cgagaatctc	ttgttttctc	gctgctgtta	gggagttcag	gaaaaagttc	960
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&lt;210&gt; 248

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-194-337 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

212

<222> 481..500  
 <223> 12-194-337.mis1, potential  
  
 <220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-194-337.mis2, potential complement  
  
 <220>  
 <221> primer\_bind  
 <222> 159..177  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 674..695  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-194-337 potential probe  
  
 <220>  
 <221> misc\_feature  
 <222> 19,208,227,235..236,361..362,690  
 <223> n=a, g, c or t

<400> 248  
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 ggtaaaattg tttcaataaa tggctaatacc agtatttttt tcttccaaaa aaacaccttt 180  
 tcccctttga tttagtgtat accacacnag agtcgcttag ccgattncag tgacnntttt 240  
 ttgatagctg gagaatgctt cctaatagggt gcagggtgtg aggggttttc ttgacatttt 300  
 cagtaagaaa gtgaatgttg tgccatagga aagctttatc acagggtgcac gttggtagcc 360  
 nnaactaaat aacgtgagtg tcaagtacag tatctagtga gtggagaata actgtaataa 420  
 ttacaattac agttgtagtt aaagtgaagt ttgtatcaaa atttctgttt caaagggtgaa 480  
 cttttaggag gtgtatctgc rgtgtttctt cctcagatgt gatttctctg aagcagtggt 540  
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 tcaaaattac tatcatatac atattttaa cactgacta acctgtgtgt ggtattatgc 660  
 aaactgctca gctcagtaag ggaacagaan ataagaaaaa aaggacaata ttgggagtc 720  
 tttatacaat tcaatttatt caattgaagg actatctttt ctttaaaaaa gttctgcttt 780  
 cttggtgtta aaataaagca tcttttatga aatgatgggtg atagtaaatg gtgatatatg 840  
 gtaaacgtaa atagtaaatg caagggtagt tattgtgatt tttaaaaata aagagagccg 900  
 agaatctctt gttttcctgc tgctgttagg gagttcagga aaaagttcct ggggcttttg 960  
 ctgttaagaa gattcaaata tagagtattt cttaaagagta a 1001

<210> 249  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens  
  
 <220>  
 <221> allele  
 <222> 501  
 <223> 12-194-479 : polymorphic base C or T  
  
 <220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-194-479.mis1, potential  
  
 <220>



213

```

<221> misc_binding
<222> 502..521
<223> 12-194-479.mis2, potential complement

<220>
<221> primer_bind
<222> 17..35
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 532..553
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-194-479 potential probe

<220>
<221> misc_feature
<222> 66,85,93..94,219..220,548
<223> n=a, g, c or t

<400> 249
gctaataccag tatTTTTtttc ctccaaaaaa acaccttttc ccttttgatt tagtgataac      60
cacacnagag tcgcttagcc gattncagtg acnntttttt gatagctgga gaatgcttcc      120
taatggttgc aggtgtgtag gggttttctt gacattttca gtaagaaagt gaatgttggt      180
ccataggaaa gctttatcac aggtgcacgt tggtagccnn aactaaataa cgtgagtgtc      240
aagtacagta tctagttagt ggagaataac tgtaataatt acaattacag ttgtagttaa      300
agtgaagttt gtatcaaaat ttctgtttca aagggtgaact tttaggaggt gtatctgcgg      360
tgtttcttcc tcagatgtga tttctctgaa gcagtgggtc tctataactg acttgacaca      420
cagtgcattc gatatttcga atgacttaag tgatagtatc aaaattacta tcatatacat      480
atttaaatac catgactaac ytgtgtgtgg tattatgcaa actgctcagc tcagtaaggg      540
aacagaanat aagaaaaaaa ggacaatatt gggagtcttt tatacaattc aatttattca      600
attgaaggac tatcttttct ttaaaaaagt tctgctttct tggtgttaaa ataaagcatc      660
ttttatgaaa tgatgggtgat agtaaatggt gatatatggt aaacgtaaat agtaaatgca      720
agggtagtta ttgtgatttt taaaaataaa gagagccgag aatctcttgt tttcctgctg      780
ctgttaggga gttcagggaa aagtccctgg ggcttttgct gttagaaga ttcaaatata      840
gagtatttct taagagtaaa tatgagttaa ttttgagatc attggtcaca gtctactgag      900
aaatccactg agaaagctgt taaatttgca tggcattaaa gatgttccta ttctgatcat      960
tgttaaatac atctttttta ttagatataa ttgtttttga g                                1001

<210> 250
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-442-133 : polymorphic base G or C

<220>
<221> misc_binding
<222> 482..500
<223> 10-442-133.mis1

<220>
<221> misc_binding
<222> 502..521
<223> 10-442-133.mis2, potential complement

```

214

<220>  
 <221> primer\_bind  
 <222> 369..386  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 777..794  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-442-133 potential probe

<400> 250  
 ggcagagttg ggaaatgcag cctgcacagg tgtgttcagg tggccggtga tatgtcacc 60  
 ccgcggccta gacttcagtc ctgcatctgt cgtgaggagt aggaagtggg gagagcagcc 120  
 cagaggccag gagctcggaa gccccagggt cccagtggcc gccctgactg cctggcctct 180  
 ccccgcaaaa ctccaggcac aatgacctcc cctggcagct gctggatatg ttcaacaacc 240  
 ggctgcagga cgacagggcc aacctgacca ccttgccggg cacacacacc aacatcccca 300  
 agctgagggc cggctttgtg ggaggccagg taccgcctgc cctgccttgt gcttgccctg 360  
 tgtggggtca tcccgtctcc tacctcaggc ctggctacag tgggaccatc cctgtggtgt 420  
 tctccagggt tccctcgggt cccaggccga gggagggctt cccagcgggt gggaaaccag 480  
 actcccacag gcatgcgggg sgtgggctga gacctggctg catcagctcc tggcaccccc 540  
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 cggaggacgc tggagcagat ggacgtggtc caccgcatgt gccggatgta cccggagacc 660  
 ttctgtatg tcaccagcag tgcagggtgg gtctcgacct gggtcctcca ggtcctgcgt 720  
 cttctcacc agccctcatc ctgagcagca ggtgccggtc aggcacctc accctccaga 780  
 taccaggtgc cactccctc gcacctgac tctccccgca ggcattcggc aggccttccg 840  
 ggaagggaaag gtggccagcc tgatcggcgt ggagggcggc cactccattg acagcagttt 900  
 gggcgtcctg cgggcactct atcagctggg catgcggtac ctgaccctca cccacagctg 960  
 caacacgccc tgggtgcgtga ctccccatgg gagggccccc g 1001

<210> 251  
 <211> 984  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 484  
 <223> 10-444-248 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 464..483  
 <223> 10-444-248.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 485..503  
 <223> 10-444-248.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 237..253  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 567..586  
 <223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 472..496  
<223> 10-444-248 potential probe

<400> 251  
gcattcggca ggccttccgg gaaggggaagg tggccagcct gatcggcgtg gagggcggcc 60  
actccattga cagcagtttg ggcgtcctgc gggcactcta tcagctgggc atgcggtacc 120  
tgaccctcac ccacagctgc aacacgccct ggtgcgtgac tcccatggg agggccccgg 180  
gctgtggtca ggaggaggg ggcagacact ccctgccacc ctccagagcc catccccctc 240  
gcctgtgagt cccaggccgg gcctcgctg ctgggctgat gggaggccga gaccaccgct 300  
cacctcttgg gcacctgcct tttgcttctc cagggtgac aactggctgg tggacacggg 360  
agacagcgag cccagagcc aaggcttgtc accctttggg caggtgagt ggggtgggagc 420  
ggccagtac ccccgaggag aaggcagagg ccctggagg tgaccagaac aatgcatctc 480  
ctcrcgtggg acctcagtgt ccttgtctgt aaaatggagc tggcagccat cccccaggg 540  
tgggtgtga gccctgagt gcccggact tccagccacg aaggatgatg actcacatct 600  
ggtccagccc gtccacctcc gcagccccga ccctgggggc tgtgaggggtg gacggagccc 660  
tgtcttccca gcgtgtggtg aaggagctga accgtctggg ggtcctcatc gacttggtc 720  
acgtgtctgt ggcacccatg aaggccaccc tgagctgtc cagagccccg gtcatttca 780  
gccactctc gccctacagc gtgtgcgcaa gccggcgcaa cgtgcctgac gacgtcctga 840  
ggctggtgt gagggccgag ggggcgacct ccacccgcc tccctgggca ggccctccca 900  
gctctcagct tcacctgtc ttccttcttg tgcagaaca gacagacagc ctggtgatgg 960  
tgaacttcta caacaattac attt 984

<210> 252  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 10-445-281 : polymorphic base C or T

<220>  
<221> misc\_binding  
<222> 482..500  
<223> 10-445-281.mis1

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 10-445-281.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 221..238  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 624..641  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 10-445-281 potential probe

<220>  
<221> misc\_feature  
<222> 926,936,957,960,972

216

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 252

```

cgggagacag cgagccccag agccaaggct tgtcaccctt tgggcagggtg agtgggggtgg      60
gagcggccag tcacccccga ggagaaggca gaggccctgg aggggtgacca gaacaatgca      120
tctcctcgcg tgggacctca gtgtccttgt ctgtaaaatg gagctggcag ccatcccccc      180
agggtgggtg ctgagccctg agtggccccg gacttccagc cacgaaggat gatgactcac      240
atctgggtcca gcccgtcac ctccgcagcc ccgaccctgg gggctgtgag ggtggacgga      300
gccctgtctt cccagcgtgt ggtgaaggag ctgaaccgtc tgggggtcct catcgacttg      360
gctcacgtgt ctgtggccac catgaaggcc accctgcagc tgtccagagc cccggtcac      420
ttcagccact cctcggccta cagcgtgtgc gcaagccggc gcaacgtgcc tgacgacgtc      480
ctgaggctgg tggtgagggc ygagggggcg acctccaccc cgcctccctg ggcaggccct      540
cccagctctc agcttcaccc tgtcttcctt cttgtgcaga aacagacaga cagcctggtg      600
atggtgaact tctacaacaa ttacatttcc tgcaccaaca aggccaacct gtcccaagtg      660
gccggtagggt ggggtgtgag cgcccaaggg ggccgaaggg ggagggcctc actcgggacc      720
catacctgct gctccctgga cagaccatct ggatcacatc aaggagggtg caggagccag      780
agccgtgggt tttgggtggg actttgatgg tgttccaagg taaggggctg agagctctgt      840
cctgtggatg agccgggagg ttcattggcc cgtcagaggg atgagggtggc tggaggaggg      900
acctgtgtcc tagtgtgggg gccangttc tcctgnctc aacacagggt ccttganggn      960
ctggaggacg tntccaagta tccagacctg atcgtgagc t                                1001

```

&lt;210&gt; 253

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-668-362 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-668-362.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-668-362.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 844..861

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 390..410

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-668-362 potential probe

&lt;400&gt; 253

```

cctgcgcctc tcgtgcacct ccacatctgg tttgaggta ggtcatttcc tgtctgtggg      60
cccagctgct gtgacacca aggggagggc cggcgctccc gaagccaggt cagccgtgca      120
cccggcagag ccccgcatat tggccccagc agagcttggg gtggggggag ctcgggtgtca      180
ccaacaggcc cttgaggaca ctcgtgtgga gaatccctgg gacacgtgga ggacccccaa      240
gtcctgagcc ccgtactccg tactgcaggg agcaggccag gagccacggg ccttgggggca      300
cagggtcctt ctcagggaca gggttcaggca ctggctggaa caggctggac ccctctaccc      360

```

217

```

agcacatgtg ggcattgcgtc tgggttccga ggggtgggaa atgtggaaa gctcctgctg 420
gccggcttag gcctgctgcc cttggagcct tccatataca gagagtccca cctccagcaa 480
gggttggcct ggactctcca wccccctgct gtgcccagga ctccccagg gacaaggcaa 540
ccagaggccc agaccctcc ccagcaaaga gaagcaccac gggagctgtc tccaagagc 600
ccttgacctg gggcccagct ggcctagggc cgggcccccg ggctgttgta gacagagtg 660
tccatctgtg cacgctgtag ccacaagcgc cgctgggctg cgccaagcag cacacgcagc 720
gagtggcctg gacggcaggc gggcagggtc gcacagtggg cagcatgcag ctggcgacag 780
gtgtcgcagc agagggtggg cagggcgcca ggggacaggc agctgtgtgc ctggtagcct 840
gggggctccg agacagacct ataggctgag gctggtaccc cctctgcccg aatacgctc 900
gggctggccg ctgcagagct ttcaggggtg gccagggtccc caggcctact cagctcacgg 960
cgcagagaga ggaaggagaa ggcggaaggt tcagggtcca g 1001

```

&lt;210&gt; 254

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-670-48 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-670-48.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-670-48.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 454..474

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 883..901

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-670-48 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 49,105,142,317

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 254

```

cttttctgga cattccatgg aaagggaact gcaacatgag tggctcttnc ctctggcttg 60
tctcactcag tgtccacact ttgggactcc ttcacgggtgt ggcangggag tcagcagcct 120
gctcctcact gtcgctcagc cngtcaactgc atggagagcg tgcacacttc attttatctg 180
ctgaaactct gttctcactc ttgctctgct gttttccctt catgtccacc ttcctccccc 240
accctctggt gtgacagaca ctgctggggc ccttccaaca ggcgggtcctc cttccccccac 300
ttccgttact ggcagangtc ctgcccactc ccattgcttca cagctccctc cagccccctgg 360
tgaatttgca ccattgaatg agtcctgcag aggcctgggca ggcgaagcct tcctgggaaa 420
ggtttccctg ctaatatgaa gcaaggagca atgcctttgc agtagatgca gtcctgcctg 480
cgtgtggggg gctgtacact satcgggggc tgtggacact gagagtgggtc aagaagagac 540
acgcaaagtc tttgatgtca ctgttgtgct aacaagccaa cctggaagct ccacctctga 600

```

218

```

cccttcattc gtgagacatt aaatccctgt gttaaagctg cgttttagtga gcttttctgt 660
aactttcagc taaaagcacc gtgacagaca ggtagcattt tccaaactgc cacccaaggg 720
tgatgctatt cctgcagggt tccatcctcc ggtggggggg tgcgccctcc tgtgaggggt 780
tccaccctcc agtgggagggt tccaaagttg ggaattcaaa tccaagtctt tcttccaagt 840
ccaaggccca gagcctccta ctttcagccc ttgatcctac aagcgtgtca gtgtgaagca 900
gtccaggccc agtcctctct cctagcaggc cagtcatccc cctgacattc tttcttctct 960
gacccgagtg gttctctctt tattattttt ttttttttct g 1001

```

&lt;210&gt; 255

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-670-91 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-670-91.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-670-91.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 411..431

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 840..858

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-670-91 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 6,62,99,274

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 255

```

tcttttncctc tggcttgtct cactcagtgt cccacctttg ggactccttc acggtgtggc 60
angggagtca gcagcctgct cctcactgtc gctcagcng tcaactgcatg gagagcgtgc 120
acacttcatt ttatctgtct aaactctgtt ctcactcttg ctctgctgtt ttcccttcat 180
gtccacccttc ctcctccacc ctctggtgtg acagacactg ctggggccct tccaacaggc 240
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ctccctccag cccctgggtga atttgcacca ttgaatgagt cctgcagagg ctgggcaggc 360
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agatgcagtc ctgcctgcgt gtggggggct gtacactcat cgggggctgt ggacactgag 480
agtgggtcaag aagagacacg yaaagtcttt gatgtcactg ttgtgctaac aagccaacct 540
ggaagctcca cctctgaccc ttcattcgtg agacattaaa tccctgtgtt aaagctgcgt 600
ttagttagct tttctgtaac tttcagctaa aagcaccgtg acagacagggt agcattttcc 660
aaactgccac ccaagggtga tgctattcct gcagggttcc atcctccggt gggggggttc 720
gccctcctgt gaggggttcc accctccagt gggaggttcc aaagttggga attcaaatec 780
aagtctttct tccaagtcca aggccagag cctcctactt tcagcccttg atcctacaag 840

```

219

```

cgtgtcagtg tgaagcagtc caggcccagc tcctctgcct agcaggccag tcatccccct    900
gacattcttt cttgtctgac ccgagtgggt ctctccttat ttttttttt tttttctgag    960
acagagtctc actctgtagc ccaggctgga gtgcagtggg g                                1001

```

&lt;210&gt; 256

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-670-157 : polymorphic base C or T.

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 12-670-157.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-670-157.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 345..365

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 774..792

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-670-157 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 33,208

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 256

```

gtcagcagcc tgctcctcac tgctcgtcag ccngtcactg catggagagc gtgcacactt    60
cattttatct gctgaaactc tgttctcact cttgctctgc tgttttcctt tcatgtccac    120
cttctcctcc caccctctgg tgtgacagac actgctgggg cccttccaac aggcgggtcct    180
ccttccccca cttccgttac tggcagangt cctgcccact cccatgcttc acagctccct    240
ccagcccctg gtgaatttgc accattgaat gagtctctgca gaggctgggc aggcgaagcc    300
ttcctgggaa aggtttcctt gctaatatga agcaaggagc aatgcctttg cagtagatgc    360
agtctgcctt gcgtgtgggg ggctgtacac tcatcggggg ctgtggacac tgagagtggg    420
caagaagaga cagcaaaagt ctttgatgtc actgttgtgc taacaagcca acctggaagc    480
tccacctctg acccttcatt ygtgagacat taaatccctg tgttaaagct gcgtttagt    540
agcttttctg taactttcag ctaaaagcac cgtgacagac aggtagcatt ttccaaactg    600
ccacccaagg gtgatgctat tcctgcaggg ttccatcctc cgggtgggggg ttgcgccttc    660
ctgtgagggg ttccaccctc cagtgggagg ttccaaagt gggaattcaa atccaagtct    720
ttcttccaag tccaaggccc agagcctcct actttcagcc cttgatccta caagcgtgtc    780
agtgtgaagc agtccaggcc cagctcctct gcctagcagg ccagtcatcc ccctgacatt    840
ctttcttgct tgacccgagt ggttctctcc ttattattt ttttttttcc tgagacagag    900
tctcactctg tagcccaggc tggagtgcag tgggtcgatc tcgctcaccg ctacttctgc    960
ctcccgggtc ccggttcaag gttcgagcaa ttctcctgcc t                                1001

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220

<210> 257  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-671-148 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-671-148.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-671-148.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 354..372  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 784..804  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-671-148 potential probe

<220>  
 <221> misc\_feature  
 <222> 31,33,37,88,270,348,764  
 <223> n=a, g, c or t

<400> 257  
 agcggcctcc ctcagctggt gtcggagtgc ncngccntgg aggagtcat ccacgagggg 60  
 tagaccctgg cgggtcaggg tgggggtncct ctgtgcgaag agcaggggacc tcatccctca 120  
 acctaaacgg tggttgctcag tagcagctgg ggaggggtagc gtgtgggtccc gtcacagcca 180  
 tgcccttcggg cagccctgtt ggggtccacct gtgcagagtg cctgctgcca catgtcccgt 240  
 ggcccttcgc ttagctcagc ctctcccagn acagcaggag ctgggggctg tcccctgagc 300  
 agcatctgcc tggggagggg ataaggatcc agtgccatcc ctggctcnta ccgtccaagg 360  
 acgagcacat agtgggtggg aaaggcctcc tctagcccag ccgcccctgac agccccacct 420  
 gcccaccctc agcaccacag tcctgaactg ggagggcaga cactaccttc tggagggccg 480  
 ggggtgtcct tctgttcgtt yagatgacac cggctgcccc tgccatccat gcaggtacct 540  
 gatcggagag gagggtact gcctcacatc actgcagagt gccctgagct acgtggagct 600  
 gctgccccgg ggaggcctgg ccaagtagta cagctagagc ccaggggtccc tgcagggcct 660  
 ggcctcgcct ccagggctg tctctcctac acctggagcc atgggatcta ctgaggacca 720  
 tcctggagcc tcacgtgct ctgcacatgg tgggggcttg tccnactgtg gtgtgctctc 780  
 cagctcttca cttcctcac ctcacatctg accctggagc ttgggggttg cactctctgc 840  
 ccgacagaat ggggtcgaat cagctccagg aggaaccagg ccctgctctc ctgtgtaggc 900  
 ctcagagagg ccaagaaggg aagctttggg cttcgggtgg tgcaggctca gcgatgaaca 960  
 tctggctggg gcagctcctg gggagcatca ggggaagagg g 1001

<210> 258  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens



221

<220>  
 <221> allele  
 <222> 253  
 <223> 12-679-245 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 233..252  
 <223> 12-679-245.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 254..273  
 <223> 12-679-245.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 9..29  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 439..458  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 241..265  
 <223> 12-679-245 potential probe

<220>  
 <221> misc\_feature  
 <222> 257..258,678,798  
 <223> n=a, g, c or t

<400> 258  
 tgcccactta cttcctttct gggtaaggct caaaggactc gtccgaagac tgggcatttt 60  
 gcttcttgct attttcatct tcaactcaaga cgtctctagg aattaagaga gaatcttcag 120  
 caaagccctg ggcctgggca gtgacagacc caacacgacc tcctgcctaa agataccaaa 180  
 tggaccccca tgtcccatat ttgagttctt ctgaaaatgt cttaaatacta aaatccccaa 240  
 atttccaatc ccrtttnncg aacacaaaagg gccagccagg ctcacaagcc agcaatcccc 300  
 acaagccatg tgctctgggc accaggacac tggacttgga aacagccctt tccacagcgg 360  
 cccagaaggg gtggaggcgc tcaaaggtag cttcctgcta agcagacttt gagaaatcga 420  
 ccacctcacc ccataactgc gaacctatg caactgatgg gtttgccatg agtcacttct 480  
 aagttcacaa aagtggctgg gtgcagaggc taactcctgt aatcccagca cttcggggagg 540  
 ctgacgtagg tggatcgcat gaggtcagga gttcaagacc agcctggcca ataccgtgaa 600  
 accccgtctc tactaaaaat acaaaaattg gccgggtgcg gtgggtcaca cctgtaatcc 660  
 cagcactttg ggaggcanga ggcgggcgga tcacgaggtc aggagatcaa gaccatcccc 720  
 gctaacaaaag tgaaaccccg tctctactaa aaatacaaaa aattagccgg gcgtggtggc 780  
 ggggtgcctgt agtcccangc gactcagaag ctgaggcccg agaattggcgt gaacccggga 840  
 ggcagagctt gcagtaggcc gagattgtgc cactgcactc cagcctgggc aacacagcga 900  
 gactccgtct caaataaaaa aaaaataaat aaatacaaaa cttagctggg tgtggtggcg 960  
 ggcgcctgta atcccagcta ctcgggaggc tgaggcggga g 1001

<210> 259  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 379

222

&lt;223&gt; 12-679-371 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 359..378

&lt;223&gt; 12-679-371.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 380..399

&lt;223&gt; 12-679-371.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 9..29

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 439..458

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 367..391

&lt;223&gt; 12-679-371 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 257..258,678,798

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 259

tgcccaactta	cttcctttct	gggtaaggct	caaaggactc	gtccgaagac	tgggcatttt	60
gcttcttgtc	attttcatct	tcactcaaga	cgtctctagg	aattaagaga	gaatcttcag	120
caaagccctg	ggcctgggca	gtgacagacc	caacacgacc	tcctgcctaa	agataccaaa	180
tggacccccca	tgtcccacat	ttgagttctt	ctgaaaatgt	cttaaatcta	aaatccccaa	240
atttccaatc	ccatttnncc	aacacaaaag	gccagccagg	ctcacaagcc	agcaatcccc	300
acaagccatg	tgctctgggc	accaggacac	tggacttgga	aacagccctt	tccacagcgg	360
cccagaaggg	gtggaggcgc	tcaaaggtag	cttcctgcta	agcagacttt	gagaaatcga	420
ccacctcacc	ccataactgc	gaacctatg	caactgatgg	gtttgccatg	agtcacttct	480
aagttcacaa	aagtggctgg	gtgcagaggc	taactcctgt	aatcccagca	cttcgggagg	540
ctgacgtagg	tggatcgcat	gaggtcagga	gttcaagacc	agcctggcca	ataccgtgaa	600
accccgctct	tactaaaaat	acaaaaattg	gccgggtgcg	gtggctcaca	cctgtaatcc	660
cagcactttg	ggaggcanga	ggcgggcgga	tcacgagggtc	aggagatcaa	gaccatcccc	720
gctaacaaaag	tgaaaccccc	tctctactaa	aaatacaaaa	aattagccgg	gcgtggtggc	780
gggtgcctgt	agtcccangc	gactcagaag	ctgaggccgg	agaatggcgt	gaacccggga	840
ggcagagctt	gcagtaggcc	gagattgtgc	cactgcactc	cagcctgggc	aacacagcga	900
gactccgtct	caaataaaaa	aaaaataaat	aaatacaaaa	cttagctggg	tgtggtggcg	960
ggcgccctgta	atcccagcta	ctcgggaggc	tgaggcgggg	g		1001

&lt;210&gt; 260

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 434

&lt;223&gt; 12-679-426 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

223

```

<222> 414..433
<223> 12-679-426.mis1, potential

<220>
<221> misc_binding
<222> 435..454
<223> 12-679-426.mis2, potential complement

<220>
<221> primer_bind
<222> 9..29
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 439..458
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 422..446
<223> 12-679-426 potential probe

<220>
<221> misc_feature
<222> 257..258,678,798
<223> n=a, g, c or t

<400> 260
tgcccaactta cttcctttct gggtaaggct caaaggactc gtccgaagac tgggcatttt      60
gcttcttgtc attttcatct tcaactcaaga cgtctctagg aattaagaga gaatcttcag      120
caaagccctg ggctggggca gtgacagacc caacacgacc tcctgcctaa agataccaaa      180
tggacccccca tgtcccacat ttgagttctt ctgaaaatgt cttaaatcta aaatccccaa      240
atttccaatc ccatttnncg aacacaaagg gccagccagg ctcacaagcc agcaatcccc      300
acaagccatg tgctctgggc accaggacac tggacttggg aacagccctt tccacagcgg      360
cccagaaggg gtggaggcgc tcaaaggtag cttcctgcta agcagacttt gagaaatcga      420
ccacctcacc ccayaactgc gaaccctatg caactgatgg gtttgccatg agtcacttct      480
aagttcacaa aagtggctgg gtgcagaggg taactcctgt aatcccagca cttcggggagg      540
ctgacgtagg tggatcgcat gaggtcagga gttcaagacc agcctggcca ataccgtgaa      600
accccgctct tactaaaaat acaaaaattg gccgggtgcg gtggctcaca cctgtaatcc      660
cagcactttg ggagggcanga ggcggggcga tcacgaggtc aggagatcaa gaccatcccc      720
gctaacaaaag tgaacccccg tctctactaa aaatacaaaa aattagccgg gcgtggtggc      780
gggtgcctgt agtcccangc gactcagaag ctgaggccgg agaatggcgt gaacccggga      840
ggcagagctt gcagtagggc gagattgtgc cactgcactc cagcctgggg aacacagcga      900
gactccgtct caaataaaaa aaaaataaat aaatacaaaa cttagctggg tgtggtggcg      960
ggcgccctgta atcccagcta ctcgggaggc tgaggcggga g                                1001

<210> 261
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 12-680-331 : polymorphic base C or T

<220>
<221> misc_binding
<222> 483..502
<223> 12-680-331.mis1, potential

<220>

```

224

<221> misc\_binding  
 <222> 504..523  
 <223> 12-680-331.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 173..192  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 645..665  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 12-680-331 potential probe

<400> 261  
 ggtaagagga tcctctaggt gggggggggg gggtcccccag gtgagggggac ctccaggtaa 60  
 gaggatcctc taggtggggg agtccccccag gtgaagggcc ctccaggtaa gagggtcctc 120  
 caggtgagag ggtcctccag cggagggggg cccccaggta agagggtcct ccagttggga 180  
 gtccttcagg tgggggtggtc ctccagggtga gagagtcttc caggtgagag ggtcctccag 240  
 gtgagaggtg aaggtgaagg tcctccaggt gagggtcagg tcaattggtc cagggatgtg 300  
 gcttcagcaa gcatttggtt gcctcctctg tgttcaggac atggggactc cgggagactg 360  
 aaggaagcaa gtggcaatgt ccatgccgtc aagttctgtc acagaaataa gcaggagtga 420  
 gtgaaagaca ggagagaagc tctgtgaggg gtggcattca agcttagctt ctgggcagga 480  
 agaggcagct atgtcagcat tcyctaacaa ggtttggcaa agtagatacc tccccctact 540  
 catggaagac ctgacttccc aggctggttag aagcttctct cacaggcacc tgcctctcca 600  
 atctctttca cagcccagcc ctggaaggtg gaaaaagaaa agttgacatg gaatgttttag 660  
 gaagcaccat tgggtccaggt ccaggatcgg ggcagccagc atgtcctcta atctggcttc 720  
 actgttgaaa tccagtcgag caagcagggc ccaaaccact gcgagaccag ctcggtcggg 780  
 cccaaaccac cgcccactgc gggaccagct cggtcggggc caaaccaccg cccactgcgg 840  
 gaccagctcg gtcgggcca aaccactgcc cactgcggga ccagctcggg cggtggagacc 900  
 ctaaccacgc ggcgctagag gaattaaaga aacacacaca gaaatatagg gtgtggagtg 960  
 ggaaatcagg ggtctcacag ccttcgaagc tgaaagcctc g 1001

<210> 262  
 <211> 384  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 154  
 <223> 10-151-154 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 134..153  
 <223> 10-151-154.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 155..173  
 <223> 10-151-154.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 1..18  
 <223> upstream amplification primer

225

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<220>
<221> primer_bind
<222> 367..384
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 142..166
<223> 10-151-154 potential probe

<220>
<221> misc_feature
<222> 346..347
<223> n=a, g, c or t

<400> 262
actcaamacc caaggagccc attctctccc ttggctttct ctcagggtcaa ggtgttgaaa      60
tgcattctcag aggtgcaggc caacaatgtg gtcctgggcc agtacgtggg gaaccccgat      120
ggagagggcg aggccaccaa aggggtacctg gacracccca cggtgccccg cgggtccacc      180
accgccactt ttgcagccgt cgtcctctat gtggagaatg agaggtggga tggtaggtga      240
tgccttcgag gccagcaag gcagaactgg gcatgccctg tgtgcgggca ctggagctcc      300
cactgagaca ctacgcact ggtccacacc ctgagagagc tggtgnnag gctgcccttt      360
ccgccacgta ggggtgccct tcat                                           384

<210> 263
<211> 426
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 205
<223> 10-138-206 : polymorphic base C or T

<220>
<221> misc_binding
<222> 186..204
<223> 10-138-206.mis1

<220>
<221> misc_binding
<222> 206..225
<223> 10-138-206.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 406..425
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 193..217
<223> 10-138-206 potential probe

<220>
<221> misc_feature
<222> 243
<223> n=a, g, c or t

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226

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<400> 263
atgatgmcca agaagccggg catgttcttc aaccccgagg agtcggagct ggacctgact      60
acggcaacag atacaagggtg ccctacagag aaggagcagt gtggaggggtg ggcggcctgg      120
gcccggggga ctccacatgg tggcaggcag tggcatcagc aagacactct ctccctcaca      180
gaacgtgaag ctccctgacg cctaygagcg cctcatcctg gacgtcttct gcgggagcca      240
gangcacttc gtgcgaggt gagggccagc tgcgggcccc tgcatacctg tgggctatgg      300
ggtggccttt gccctccctc cctgtgtgcc accggcctcc caagccatac tatgtcccct      360
cagcgacgag ctccgtgagg cctggcgat tttcacccca ctgctgcacc agattgagct      420
ggagag                                           426

<210> 264
<211> 426
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 351
<223> 10-138-352 : polymorphic base C or T

<220>
<221> misc_binding
<222> 332..350
<223> 10-138-352.mis1

<220>
<221> misc_binding
<222> 352..371
<223> 10-138-352.mis2, potential complement

<220>
<221> primer_bind
<222> 1..18
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 406..425
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 339..363
<223> 10-138-352 potential probe

<220>
<221> misc_feature
<222> 243
<223> n=a, g, c or t

<400> 264
atgatgmcca agaagccggg catgttcttc aaccccgagg agtcggagct ggacctgact      60
acggcaacag atacaagggtg ccctacagag aaggagcagt gtggaggggtg ggcggcctgg      120
gcccggggga ctccacatgg tggcaggcag tggcatcagc aagacactct ctccctcaca      180
gaacgtgaag ctccctgacg cctacgagcg cctcatcctg gacgtcttct gcgggagcca      240
gangcacttc gtgcgaggt gagggccagc tgcgggcccc tgcatacctg tgggctatgg      300
ggtggccttt gccctccctc cctgtgtgcc accggcctcc caagccatac yatgtcccct      360
cagcgacgag ctccgtgagg cctggcgat tttcacccca ctgctgcacc agattgagct      420
ggagag                                           426

<210> 265
<211> 1001

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227

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-586-414 : polymorphic base A or G

<220>  
<221> misc\_binding  
<222> 482..500  
<223> 12-586-414.mis1

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-586-414.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 88..107  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 552..572  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-586-414 potential probe

<400> 265  
gtggtcagct atcccaaggc tcttggtgt tgacatttta tgttcagctt gcacttctct 60  
ttctctcttt tagtgctgc aacttctcac ccacacttcc aacagagaat tcagccaggt 120  
gcacggcagc gtcagtgc gtaagggtgag cacattgact gtaattttta gccagtatgt 180  
tgataactga ttctccaca gcagcccaga ttacctatct ctgggttttg ctgcttttaa 240  
gcaacttgct atgggcatag cattgtattt gaaaatttat gacatactgc tctgggtattc 300  
attctaattt ttcagagtcc gaacactgac ttctgaagat aaaagtactc ctttgtgtct 360  
cttagagtga ttatcagatg ggaaacattt tggctttttc atgactcctt tggaggagaa 420  
tattctatgg ggaggtggta tgttattctt tgccagggtg caaggaaacc ctgaggttcc 480  
tgggtggcata aagttttatt racttcaaca aagagtgaag taaacacttc agagaatttc 540  
tgtgttattc actcaattct agtcagcttg accttaaacc ctcccggctc attcctgcct 600  
tgggggtcttt gcacacctac tatttctttg cctggaattg cctttctcca ggtattgcc 660  
tggttggctc cttacctct ttcatctttc tactccatta tcacctctg tgagtcgact 720  
ctgttaacac ctgataataa atctgacagc tgtctgaaac tttctaccta cccctctttc 780  
ctgctttatt ttttgcata gcactaatta ctacctgatg ctttttaaag tgtttgttga 840  
gtgggttatt gtctgtctta agctccaggg aggcagatcc tctgccatat tattagaaca 900  
atgcctagta catggaaggc gcaccactgt aataccactt atattaattc aggggctggg 960  
tgtgggtggtg catacctgta atcccagcac tttgggaggc c 1001

<210> 266  
<211> 999  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 499  
<223> 12-587-379 : polymorphic base C or A

<220>

228

<221> misc\_binding  
 <222> 479..498  
 <223> 12-587-379.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 500..519  
 <223> 12-587-379.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 857..877  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 478..498  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 487..511  
 <223> 12-587-379 potential probe

<400> 266  
 cagcactttt ctactctctc cggtcccat ctccattgct ctgtactctt ttcttttttc 60  
 ttgtgctgag aatctcgta gtagcatgtg gcctaacaaa aggaaaaaat gtttttaaac 120  
 acacacacac acacacacac acacacacac atacacagac aaaaacacaa aaactctgag 180  
 gggatctggt gaatctccaa attattgtgg gtgtactttg gcttcctttt gtatgatagg 240  
 tccccatcat gaccacctct gatgtctgtg ctgctgtcac caggcacctt tgtttttcaa 300  
 gacaacatac ttttttttcc ttttctctgt ttgtgatata actttaattt ttcttgggtg 360  
 gcttagagac taaggaggga gacatctggc ctttttagaa cctgagagga aaaaaagagt 420  
 ctttttttcc cctctgtctc tttttgcat ggctaattcc tgcatttcca ttcagggaaa 480  
 aggtggtagt gagcatagma ctgcaacagt tatattctga gtcaaagttg gggcttttta 540  
 cggcataatt atggaatttt tatttactgg tagagaggag acgagaggct ttttcagtgg 600  
 gcctgggaca gtggctgctc ttgactttgt gtgaaggga atgccaagga tgcttctggt 660  
 ggacttcagg ggacccagg gtttgccgt gggccgtgat ggcagcaggc ggtgggatgc 720  
 ttgtagctcc tcacagcagg attcctgcc actgtttttt ctctgttggg agggaagctc 780  
 ttttctagga gtgtctcagt tctgctttg gcattagtga tgggtggtgg acagttggaa 840  
 ttagtgccat gtcatacaca aatgttcac aaggcgggag tgtttctact tctggtgata 900  
 aacttgatgg tcattgttat gattaagata atgccgggca ggccgggcac agtggctcac 960  
 gcctgtaatc caagcacttg gggaggccga ggcgggcag 999

<210> 267  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-588-103 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-588-103.mis1, potential complement

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-588-103.mis2



229

<220>  
 <221> primer\_bind  
 <222> 585..603  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 60..77  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-588-103 potential probe

<400> 267  
 gcttaatgaa ttcattgtttt gttttcaaat gtgtccgtgc tctgtttttt ttatcctttc 60  
 ttttaggtcg tatttcggaat aagcctgagg tggatgaagc tgcagttgat gccatcctct 120  
 ccctaaatat tatttctgcc aagtacctga agtcttccca caactctagc aggtgggaca 180  
 cccagagcag tgtgaagaag tccacacttg caggcgtaa ttggtacacc gttaggtgcc 240  
 ttattcatta atgactccca gttcggacaa agaaattaac tcccttctcc cttctagctt 300  
 caaaaatctc tttatttctt cacctgcctg ctgtactctc caaaaagaaa gaaagaaagt 360  
 attgcagata tttgtatgtg atcagttact cttagagaat ggaagtgatc ctgtcccatg 420  
 tgaagtttga atagatgtaa caagtataa atgaaaattg gagaaagaaa acagtatatt 480  
 cccaaggat ttaaaagtac raattaatta ttgccaagat taaatttttt cctgtgaaat 540  
 ggttgctgtg gagagaaagg tttctctcac tccccaagta tcatggaaat gtgccctctg 600  
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 aattatcttt tataatcagg tgataacaga cagttcgtaa agtacatagc cattctgctt 840  
 tcctttgtaa gactagaact aaaccagctg ggatcctctt acctcaccca ggaaggaggat 900  
 tctcattgtc tcagaggctc cgtctctata tcctaggggc tgggaaacag gagcggctgc 960  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 12-589-152 : polymorphic base T or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-589-152.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-589-152.mis2, potential complement

<220>  
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 <222> 636..654  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 190..210  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-589-152 potential probe

<220>  
 <221> misc\_feature  
 <222> 393,614,617,695,819,849  
 <223> n=a, g, c or t

<400> 268  
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 aaaaaaaaaa agacatcaaa ggcattgaggt gtaggcacat agtgacagaga ggactccagg 180  
 caggggacat gtgcgatggc tctttaatgg cccaggcctt gaaatgggca agtctagctg 240  
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 catttgatac cttacgcca attccttctc ttttaaatc tggtgcagta gttgtgtgtt 360  
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 ttcaactctc tgtttcttag catttctctc tctcctgctt tggagcagac tgaaagggaa 480  
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 cttgggtttg tttgatttat tgatacaagt gtctgtcttt gtcttttctc ttatgttcga 780  
 agatcagcat ataccaaatg ggaagactgg aatcatagnt aggaactaga ttctatttcc 840  
 ctttactna attggaatga acactctagc tttgtgacca ccagttttta ttgctttttt 900  
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<210> 269  
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 <212> DNA  
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<220>  
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 <222> 501  
 <223> 12-592-118 : polymorphic base A or T

<220>  
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 <222> 482..500  
 <223> 12-592-118.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-592-118.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 384..402  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 830..849  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513

231

&lt;223&gt; 12-592-118 potential probe

&lt;400&gt; 269

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cccttgagcc caggagtttg agactgcagt gagctatgat ggtgccactg gcgcctgggt    180
aacaagcaa gaccctgtct caaaaaaaaaa aaaaaaaaaa aaaaggagct gggcatagtg    240
gcatgtgcct gtaaatgaaa ggcattcattt catgcaagct cctctgagcc caggagtcc    300
agcctagccc aggcaacagg gcaagaccac gtctcaaaaa aaaaaaaagt actctttcca    360
cctaaaatat attcaggtca tttgagttca gtttgagttc agctacgaga attatttagt    420
tgagtgtagt tcagagctgg gattttcaaa tctgccctta ctggtatgtt gctttacaca    480
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tgtagaactt gtgctcatgc tttgatttgg gatttgggga gttagggcaa gccagaaagt    600
ttttctgggt gataataatg tgggttgact ttcttaagca ttttaagcca agcacttgag    660
tttctaacia ctaaaaagct aagtcagcct gacacagctc tagcgcgccc tggcttgatt    720
ctgttcattc ccagggggag acttgccctt gttccagtc tgctctccca agccagctta    780
ctgtagtatt ccagcaattc tgagaagcag tattttttac tgctgattag aaccttaaca    840
tggaatgga aagtttgtgt agcatgtaac attattagaa gggaaaacat gactatatga    900
ttagatacat attgatttta tgcaatatgt ttgcataaaa tctcttcagt agtaacttgg    960
ttaattatt cacattgact cagcataatt tctcccttgc t                                1001

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&lt;210&gt; 270

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-593-174 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-593-174.mis1, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-593-174.mis2, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 658..675

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 138..158

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-593-174 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 22,187,214,652,684

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 270

```

agggtcaaagt ataaggattc tncaaaaatg cagagtaatt atgaaaattc tgtttaaaaa    60

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232

```

gcttcagaat agaatgttta tatagcattg atttggagct aagggtata ttctggtaac 120
ataacaaggt gctatctagt gaatgtgtga ttgtgtagtc atggagctga tgtccctgcc 180
atagaantta cagaattatc taaaaaggga aatnctataa taatggcctt caatgctccc 240
acacttggac tagccattgc cataggcaga aaggcaatct catctttact gtgtgtgcaa 300
acagtacttt aatgtaatgt gtgcttaata taagctttct ttaaaaaaaa aaaaggtggc 360
tcctgttttt gaatagctat ttttaagata gatatagtta ggaatctaaa tgtgttctat 420
atagttaata tccattatga ggtggctctg aaaaatcaac ctagtataag ttggatggct 480
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aaaaaaaaa aaaagacatc aaaggcatga ggtgtaggca t 1001

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&lt;210&gt; 271

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-596-124 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-596-124.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-596-124.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 378..397

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 805..825

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-596-124 potential probe

&lt;400&gt; 271

```

gacagaccag gtcacttggc ttccgagatc atcagagaag ataagtctgt ctctttcagc 60
tgccagtaag ttttccagga tgagagggga aaaagaaagc ctccagtgc ttcagtgtgt 120
ttgccagtgt tcttgggatt gttttacacc atcctttact tcccttgcgc agacctctct 180
gtttcaccat tgctcaggca ttcaggaaaag tatctgctca ctcccacttg gtgagtcttc 240
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caaaggccag gggtctgcgc gcctctgcag agttgtgtgt agggagactt gtgtcatcat 360
ccacaacctt gtttctcact tcctgggttg gctcatctct gaagaacagg tctcccagct 420
tcgctcctta tctactgcatt gtgaagagga ggaaaagtga atcacggaga gagaaaggaa 480
aggatagaat cacaggctgc rtctgcacct gaaaagtgc cgcgggaaac tctatggcgg 540
attttttttt taactttctt cttcctgtta aaacataggt cactaactgt gatgttattt 600

```

233

```

gttttctaag tggatatgtga gatttttctaa tgtagttaga agtttcattg tctgatggac 660
acaatatgcc cttccggttc tattcaaacc agcaggatct gtcggtgctt agagatggct 720
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tagaacaggc caaactggac tgtctgttca tagcgtgcct g 1001

```

&lt;210&gt; 272

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-602-196 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-602-196.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-602-196.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 307..325

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 704..724

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-602-196 potential probe

&lt;400&gt; 272

```

ggccagcctg ggtttctcca gttgggtctc ataatgcac tgcaccctgg cgcagcatgt 60
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gtctctgtag taactttctt gtctacctgc atttttcttt cagacaccaa acacctttgc 660
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&lt;210&gt; 273

234

<211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-602-350 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-602-350.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-602-350.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 153..171  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 550..570  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-602-350 potential probe

<400> 273  
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 tcaaggagcc tctttacagt aactgggcta aacattttgt tgcgtccgt cggccttatg 180  
 tcttcatcta taacagtgtgac aaagaccctg tggagcgtgg aatcattaac ctgtccacag 240  
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 aatatggcaa ggcagtcctc attgctgtct ctgtagtaac tttcttgtct acctgcattt 480  
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 acaatacggg aagaagtgtt gttgtgtgtg ttgtgtgtt tgagacggag tctcactttt 660  
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 cgattctcct gactcagcct cccagtagc tgagactaca ggcacgtgcc accattgcct 780  
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<210> 274  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-603-191 : polymorphic base T or C

235

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<220>
<221> misc_binding
<222> 502..520
<223> 12-603-191.mis1, complement

<220>
<221> misc_binding
<222> 481..500
<223> 12-603-191.mis2, potential

<220>
<221> primer_bind
<222> 668..688
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 240..260
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-603-191 potential probe

<220>
<221> misc_feature
<222> 339
<223> n=a, g, c or t

<400> 274
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aattttgtag tccctcctccc atgtaaacct actgtaccta ggatagagta gacctgttc      180
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ctgagtccat tgcagtcttg caaaactggt cactttcttc tctgttttgt cctgtgtggt      480
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aaggcggttg gatcacgagg tcaggagttc aagaccagcc tggccaagat ggtgaaaccc      840
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tactcaggag gctgaggcag aggatcgctt gaaccggga ggcggagggt gcagtgaacc      960
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<210> 275
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-783-73 : polymorphic base G or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-783-73.mis1, potential

```

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<220>
<221> misc_binding
<222> 502..521
<223> 12-783-73.mis2, potential complement

<220>
<221> primer_bind
<222> 429..446
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 858..878
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-783-73 potential probe

<220>
<221> misc_feature
<222> 202,209,236,281,447,461,627,992,995
<223> n=a, g, c or t

<400> 275
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tttttattca tgtcatcctc acttactttg actattgttt gtttctactgg attgctgtta      180
attgttcttt ctaaaacatt anccttagnt tttctttcac agaaacagtg ctatttcttt      240
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agtgcctttac tatatgccaa gcatgattct aaatgctggg gatatactag tgaagaaagc      360
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gcagagcctg ctggcaaccg tgagcattgg tcagcgtgga cattggacaa ggggcttctt      720
tgctcagccc tcattcctct aacatggttc tctcctgtgt tctgcatgta ggcctttgag      780
gattggaata agacagagct agactcattc ctgattgaaa tcacagccaa tattctcaag      840
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<210> 276
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-783-421 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-783-421.mis1, potential

<220>
<221> misc_binding
<222> 502..521

```



237

&lt;223&gt; 12-783-421.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 81..98

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 510..530

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-783-421 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 99,113,279,644,647

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 276

agtgaagaaa	gcaaagtgcc	tgccctcccg	gagctgggaa	gacagacaga	aataaatata	60
caatatcaaa	ctagtttagcg	atgactgtga	gaacgaagna	gtaaactggg	acnaagatcc	120
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&lt;210&gt; 277

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-785-200 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-785-200.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-785-200.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

238

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<222> 302..322
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 791..811
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-785-200 potential probe

<400> 277
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ctaatacaggc aactgatacg ttttatggaa aacatctatt tggttaactg atgtctgaag      180
tgacaataat agtgagttga cctttgtcct acaatctcct aaaaagagca gcatctctac      240
ctggtgttag ctctggggaa aagggcatgt tgcttctgtt gaaagcagcc tcaaaggcgt      300
cctgtgtttt tatagtcgct cctctggcct gtgccgggaa ggtatggctg cagagcactg      360
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<210> 278
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-785-393 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-785-393.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-785-393.mis2, potential complement

<220>
<221> primer_bind
<222> 109..129
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 598..618
<223> downstream amplification primer, complement

<220>

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239

<221> misc\_binding  
 <222> 489..513  
 <223> 12-785-393 potential probe

<400> 278  
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 ctgtcgtcag ggaacccag gaatgctgcg gtgccagcaa tacaggaatc agggatttgg 240  
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 ggttcctctt taactgtgca gttcttttta ttccagacta ataataggga gagagagaga 480  
 gagaacgata acatctcact rtctgtgaaa ataactaaat atagccatt gttttaacaa 540  
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 ctagtaatgt gggaacacaa agctgccagg gtaagtcagg aaagactgcc tggaagaggt 720  
 gacatttgaa tccacaacag actcttatgt aactatgccca tccacaggct ctgcactaa 780  
 ctgtggaagc ttgagtctgc taaattttgt ctgtacaaaa tgacattgtt tttattttta 840  
 gtaaaaagct tttacaaagg aagaggagtt aattgaatgt acttcttcac tctctctaaa 900  
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<210> 279  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-787-103 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-787-103.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-787-103.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 583..602  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 74..94  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-787-103 potential probe

<400> 279  
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 gagaagactc cttctgtgta gacccctctt gagcctgtgg actggacatt tgccagagg 180  
 gagcttctgg aaaaacaagg aattgatatg aaacaagaga tggagaaaag gtaatgcaca 240

240

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aaggaagaag cagatcttct tttggagcag cagagactgg taggagtcct gaatctgcta 420
aactgttggg aaaagggcag cttgttccca tactttccct gtccacaga gcagtactca 480
cccaaattgc ttctgtctca rtgataccaa gcactattct ttaatttcct taatggagaa 540
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&lt;210&gt; 280

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-790-396 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-790-396.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-790-396.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 876..896

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 423..443

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-790-396 potential probe

&lt;400&gt; 280

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tgcaagtggc agattgtagc tcgctgcagc ctcagacacc tgggctcgtg tgattctcct 180
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cctcccgagg cctgtatttt tttatgtatg agtaggtttc attactttag tctggaactc 420
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tcacttgagc ctttttcttc atctttaata ggtactataa agctttatac aaatgtatta 600
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acttatgaga agtagaacat atgagaaatg acaagaacaa attttctttt tggatctaga 720
tatgcagatc gtgcaaaaca aattaaatgc aatgctgtta tcaatgagga cccaatgcc 780

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241

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cttcttcagg gttcttattc agcgttctta tatttaaaat aaacttcaag ttaaggagca      960
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&lt;210&gt; 281

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-791-211 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-791-211.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-791-211.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 291..311

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 671..690

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-791-211 potential probe

&lt;400&gt; 281

```

tacattggaa tcttttagtt tgtaaattgca gaaatgattc tgaacctgta ggataccact      60
attaaagaca gacatgtttt agaactgccca tctagaacag ttgtaacctg tgtttgtatt      120
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gtcttttttt ctctctctgc ttctttccct tttccctact tttcctgcct tctcttcttt      240
ctatctccca gatctgaaa agtttcagaa caataagcat agatacttgc tagcctctga      300
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ccgaggcggg gtggatcacg aggtcaggag atcaagacca tcctggctaa cacggtgaga      960
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&lt;210&gt; 282

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-792-233 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-792-233.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-792-233.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 712..732  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 284..304  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-792-233 potential probe

<220>  
<221> misc\_feature  
<222> 204,218,454,485..486,512,566,576,579,614  
<223> n=a, g, c or t

<400> 282  
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gagaaaatggg cataatttaa accattatta tattgttgag gtatccctag ctattattat 180  
agcaaagtgg gaaaaaagtg ttnatttcta ttgaagtnta tgtaatgac cgacattaat 240  
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tcctttgtag attngcctta atgatttgta caaatgactg ggaggcgggg atgctgcctg 660  
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ctttcccaga ttttaaaatc tgttctagat attcttagct tgaaccactt ttgattgtga 840  
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<210> 283  
<211> 989  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 505

243

&lt;223&gt; 12-793-383 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 485..504

&lt;223&gt; 12-793-383.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 506..525

&lt;223&gt; 12-793-383.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 866..884

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 365..385

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 493..517

&lt;223&gt; 12-793-383 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 505

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 283

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&lt;210&gt; 284

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-803-125 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

244

&lt;222&gt; 481..500

&lt;223&gt; 12-803-125.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-803-125.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 605..625

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 169..189

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-803-125 potential probe

&lt;400&gt; 284

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&lt;210&gt; 285

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-805-115 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-805-115.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-805-115.mis2, potential complement

&lt;220&gt;



245

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<221> primer_bind
<222> 596..615
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 135..155
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-805-115 potential probe

<220>
<221> misc_feature
<222> 313,721
<223> n=a, g, c or t

<400> 285
gagccaccgc gcccggttc tgtatgtgat ttctttttca ctctaattca ctttactaat      60
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aaaaaacaag ggtaatttgc gtgtgctgta acaccactt cttggaccct atgtggagga      180
tctgtccaag ttggcagtta ctccctacac agacattgct gacctcatgg atgctgggaa      240
caaagccagg tatggttagga aatagagtaa tgactgaggt ctttggcacc ttttgaggtc      300
cttttttccc agnttaaggg tttgaggcca catttatagc tatgaaagtt gctttaattg      360
tggagtccctc tgatcctttg actgtgctgt aacagtgagg gcttcagagt gttaagtatc      420
ttctcatcta aacagtacat tgctgaaaca tgttagagct ctcattgagt ttactctta      480
ggttttgatc aagtcatggt raaatgttag ttatcatgta tatatgcggc tgtggtagta      540
attgattctg gaataaatga gcaccaataa agaataaaaa ttaaatgctt cattggtttc      600
taaaagggata gctggagtat gaaaaatgat tctgatgggg tgcctttgga attggagagt      660
aacttgatgg gctcacaaat aaggcagtat tgagagcaat tgtgtaagtg acaatcttgc      720
naatctgtga ataagtttaa gactattgct tgtgattggt atgcttataa tgtggagcct      780
gctgtccatt tcaactctca tctaagaatt cccttgtcac gtggtcacct ttatgtttat      840
ttaggacagt ggcagctaca aacatgaatg aaacaagtag ccgttccac gctgtgttta      900
cgattgtttt caccagaag aaacacgata atgagaccaa cctttccact gagaaggtag      960
gagagtttca gtctctaggc ttgagttgtg aaggatggag a                                1001

<210> 286
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-808-52 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-808-52.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-808-52.mis2, potential complement

<220>
<221> primer_bind
<222> 450..469
<223> upstream amplification primer

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246

<220>  
<221> primer\_bind  
<222> 894..914  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-808-52 potential probe

<400> 286  
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gcccaccta taattatattt tttagatgg agtttgactc ttgtcaccca tgctgggtgtg 120  
caatggcacc atctcagctc actgcaacct ctgcctcccc agttcaaaca attctctttc 180  
ctcagcctcc caggtagttg caattacagg tgcccgccac cacacctggc taatttttaa 240  
aaatattttt agtagagacg gggtttcagc acattggcca gactgggtctc aaactcctga 300  
cctcagggtga tccgcccacc tcggcctccc aaagtgtgtg gattacaggc gtgagccacc 360  
gtgcctggcc atgttgggtt tttttgggtg ggggaagggt aatggggatt tgaaaagttg 420  
aacatgtcaa tttaatttaa caagccctca tcacgagcag agcatgagca ggggccactg 480  
tcctgggtgt ttggggagaa rcagaagaag ggagggagga gcagccctca ggttaaaagc 540  
aatgatttgt tccaaccaat atttgagtac ccacctgtg cctggtgtgc tgctgggtgc 600  
agtaattcag cagccaacaa agcaagccca tgcactgtg ccctccatt ccagtgggga 660  
agacaatgtt taaacaaatt tacaacatcg gtcctaagta ttttggagga aagcagctaa 720  
ataaaaggat gaaagaggga tgggaacgag ggatgtttag atggcatagt cagggaaggc 780  
ctctcttcgg agatcacatt tgagcagact ctagaatgaa aggatagggt aagccatgca 840  
aaccctggg caaagaggtc tccaagtaga gaaatgaagc ccttgaggca gcagtgtcct 900  
cagtaggttc cagaaatagc aaggctcgtg gtacagagtc agggactggg agcgcaatag 960  
aagacatcag aaagtagcca gggaccagga tatttagggc t 1001

<210> 287  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-808-75 : polymorphic base G or C

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-808-75.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-808-75.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 427..446  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 871..891  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-808-75 potential probe

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<400> 287
ctcccaaagt gctaggatta caggcttgag ccaccgtgcc caacctataa ttattttttt    60
gagatggagt ttgactcttg tcacccatgc tgggtgtgcaa tggcaccatc tcagctcact    120
gcaacctctg cctccccagt tcaaacaatt ctctttcctc agcctcccag gtagttgcaa    180
ttacagggtgc ccgccaccac acctggctaa tttttaaaaa tatttttagt agagacgggg    240
tttcagcaca ttggccagac tgggtctcaa ctcctgacct cagggtgatcc gcccacctcg    300
gcctcccaaa gtgctgggat tacaggcgtg agccaccgtg cctggccatg ttgggttttt    360
ttgggtgggg aagggtaaat ggggatttga aaagtgaac atgtcaattt aatttaacaa    420
gccctcatca cgagcagagc atgagcaggg gccactgtcc tgggtgtttg gggagaaaca    480
gaagaaggga gggaggagca scctcaggt taaaagcaat gatttgttcc aaccaatatt    540
tgagtaccca ccctgtgcct ggtgtgctgc tgggtgcagt aattcagcag ccaacaaagc    600
aagcccatgc acttggtgcc tccattcca gtggggaaga caatgtttaa acaaatttac    660
aacatcggtc ctaagtattt tggaggaaag cagctaaata aaaggatgaa agagggatgg    720
gaacgaggga tgtttagatg gcatagtcag ggaaggcctc tcttcggaga tcacatttga    780
gcagactcta gaatgaaagg atagggttaag ccatgcaaac cctgggcaa agaggtctcc    840
aagtagagaa atgaagccct tgaggcagca gtgtcctcag taggttccag aaatagcaag    900
gtcgttggtg cagagtcagg gactgggagc gcaatagaag acatcagaaa gtagccaggg    960
accaggatat ttagggctta ggtgaggact tggattttca t                                1001

<210> 288
<211> 987
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-809-119 : polymorphic base G or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-809-119.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-809-119.mis2, potential complement

<220>
<221> primer_bind
<222> 383..402
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 888..908
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-809-119 potential probe

<220>
<221> misc_feature
<222> 511
<223> n=a, g, c or t

<400> 288
gtggagtga gttcaaagaa gtatttttat atgtggactt gacctttggt cttttattct    60
cattttccac ttaagaaaat cttggctgtg tgaatgaaag agtgatactt ttaaggttat    120

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248

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agaaagtgaa atgtaatcat gccagataat tttatataga tatttttatg tatggctgac 180
ctggatgact ctaacagtg c atgtgtttgt gagtgtgtgt gtgtgcatgt gcgtgtattt 240
aatgagaaaa gtaaacttgt gtataggagg cttaaaaaat gtgtaggga ttttaggtga 300
ctgttctgat tccagacact tttattatgg aagcaatcaa gtaagtatag gaagaaatat 360
taataaaaagg ttattttattt ctctttttac tctttacagc ccgaagatcc ctgttttgca 420
tctcaaaacc gtgtgtacaa tgacattggc aaggaaatgc tcttacacgc ctttgaggga 480
tataatgtct gtatttttgc statgggcag nactggtgct ggaaaatctt atacaatgat 540
gggtaaaaca gaagaaagcc aggcctggcat cattccacag gtgaaaaaca aaacaaaaca 600
aaaatcttct cttcattatt agtggttagtc ttaaattgct ttaacagtta tttttatttg 660
gcgaacattt atgcggggat tgttttatgt caggcacaaa gatgaacaac ccattatttt 720
ccctcagagg agctcacaat tgaatgggaa ggattgacat gtacacatgt ctgtcattaa 780
aggtgggaaa gtcagtgttt tgtaatgatt ttgccatata tccaatgcca tattattttg 840
tcatttgaaa agtgttacca gcttggttaa gctctgttct aagtcctgga gatgggggga 900
tattgttgat ctgatttttt ttcaaattcc atgcatagat tacctctgaa gaatgtgatt 960
atttttgggt tttttgatag ccttatt 987

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&lt;210&gt; 289

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-810-77 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-810-77.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-810-77.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 558..577

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 126..146

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-810-77 potential probe

&lt;400&gt; 289

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ttacaagcat gagccacat gccaggtctt cctgtcaaat aagggttttaa acatgcagct 120
gtagccacag ctctcgtgtt tattagtctg gcatcacagt cagtactgtg agtgccgttg 180
cttaggctta gcctataaat atccacaaat atccattaca gaagcaggca agggcaaatg 240
ctgtgctaag cagcccaagg gccacatttg gaaccacatt aggaagcatg ctatgttcag 300
tacagtaaaa ccttggtgat tcaatggctt caacaaatgt tctgtgtagt aagtgttttt 360
catggaaata tttctgactt gtcctggtga tttctgtgtt tccttaactg aatcatcatc 420
aagaagagat taaattgttt agctatatga acataattta ttaaccagtg ctatacaaat 480
aactagagat gcagacagat rtgacgtgag accaaagagt gaatgacacc tgagcctggc 540
attcctcgga gcttaaagaa gagttcctgg tgccctctg ttttttgga tttttttttt 600
ttcaaagaaa gggaggagga ttaaagattt atctcctcaa cccctcattt ctgaaccatg 660

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249

aatccacatt	tcaagttcat	cttacattag	gcccctagag	tgggggatgg	gaagccctgg	720
acaccagcat	gcctgcctgc	cagcagaatg	agttgatgct	cctgagactg	aatgctgttt	780
tgcacttgge	ttccctatatt	atactaagtt	tgctactcag	gcttgaacag	aatatccttc	840
actttttctt	aacttagggg	gaactgtttc	atagaacatt	tacttgaaag	cctattgatc	900
ctgttttatt	tgaagaaaga	acagtggagtc	agcaaatagc	tctagacctg	tctcagtgag	960
tagtgtgggc	tcacttagct	tctctgggtc	gttggaaacac	a		1001

&lt;210&gt; 290

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-265-178 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-265-178.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-265-178.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 324..341

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 662..681

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-265-178 potential probe

&lt;400&gt; 290

ctgggggtcg	cctgagctca	cttgggggtc	tgtgacctg	gccctacggc	gtctcggggc	60
cagagctcct	tccctgcggg	cccggccccc	tgccctctcg	gccgcgcagg	catggggcgg	120
ggcgggtccc	ttcgagggcc	agggaggagg	cgcgggccagg	cctcgccgag	tctgcaggga	180
cacctgcggg	acgcgtgcca	gcgggagccg	ggcgcgaggg	cggggctggg	ggccgcctgc	240
ccggccgagg	ctctgcagcg	tgcgcgcccc	gcttctgggg	gcccgcggga	ggcgcgagg	300
aaagtgcaga	ctcccagtca	cggccaaatg	tgggaaggacc	ggacccctgg	gttgcagcgc	360
gtcgagcggg	gctgactctt	tcctttgttc	tgtttctgcc	tctctagagc	tgacatcgcg	420
ctgatcggat	tggccgtcat	gggccagaac	ttaattctga	acatgaatga	ccacggcctt	480
gtggtaagcg	gcgtgggcgc	rttgtcttct	ctctgggtcc	cgggcgcttt	agccgaggcc	540
ggcgataagg	ttgggagctt	acgggtctcc	tggccgtgct	ttgctaattg	gctctgttgc	600
tgctcgtggc	atthttgtat	ggaaaggaga	agcaccctgt	aggcgtgggc	gggccgatcc	660
cgaacttagt	cctgcggagt	gtgcctgtgg	gtccgtgagg	ttcacagccc	gaatgaacga	720
attagtgtct	taagttagaga	agaagggtgc	gggaggagaa	cccttgctgg	ctgtgtagag	780
ctctaacttg	attgcctgat	gagatctggg	gagagtgaag	atgtttcttc	taaatgttaa	840
agtggccttag	acgccaggat	gatcaataac	aacagataga	gcctgttgaa	ccggccagtt	900
cctggatttg	atthttgagtc	ctaacacggt	gggtgtggat	tctacctgac	acaccggggg	960
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&lt;210&gt; 291

&lt;211&gt; 1001

250

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 503  
<223> 10-266-203 : polymorphic base C or T

<220>  
<221> misc\_binding  
<222> 483..502  
<223> 10-266-203.mis1, potential

<220>  
<221> misc\_binding  
<222> 504..523  
<223> 10-266-203.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 301..320  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 701..720  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 491..515  
<223> 10-266-203 potential probe

<400> 291  
cagcccgaat gaacgaatta gtgtcttaag tagagaagaa ggggtgcggga ggagaaccct 60  
tgctggctgt gtagagctct aacttgattg cctgatgaga tctggggaga gtgaaaatgt 120  
ttctttctaaa tgttaaagtg gcttagacgc caggatgatc aataacaaca gatagagcct 180  
gttgaaccgg ccagttcctg gatttgattt tgagtcctaa cacgttggtt gtggattcta 240  
cctgacacac cgggggtagt tggccttcgc ctcggttgtc cgccgttttc atcctactga 300  
ggtagacataa ctttacagtg agcctccac agctggggga agaaaaggca aaggcaggtc 360  
tgacctcccc agaacttggg ttgaatggaa aggtcattgt tactttggcc acagtctgaa 420  
agtcttggtg gtcttggtgc tcctactcag gacttttgtc cttctaggtc tgtgctttta 480  
ataggactgt ctccaaagt gaygatttct tggccaatga ggcaaaggga accaaagtgg 540  
tgggtgcccc gtccctgaaa gagatgggtc ccaagctgaa gaagccccgg cggatcatcc 600  
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tgctctcagc tgctaccacg atagcagctg tttttgggtt cttcctttag ttctccttct 720  
tttaactcta gagatttttt tttttcaatt tctgctaagc tctgaccaa tgattgctta 780  
actgttgagc tgttggttaa cttgataagc gcttatgaag taactttgtc ctttgggtgt 840  
agtaattctg agaatactaa cagacatttc aataaacatt aaggcggggc acagtggctc 900  
aggcctgtaa tcccagcact ttttgggagg ctggttcaga tggatcggtt gagcccagga 960  
gttcgagacc agcctgggca acatggcgaa accctgtctc t 1001

<210> 292  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 10-403-312 : polymorphic base C or T

<220>

251

<221> misc\_binding  
 <222> 481..500  
 <223> 10-403-312.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-403-312.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 190..208  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 593..611  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-403-312 potential probe

<400> 292  
 cccttttcctt ccttgccagcc ccatcggtggc ccagtacctg tctctgcccc ctgtattctt 60  
 cttgcatgca ctgccatgca gcctggaatt tgaggctacc cagtgcccc acccattctc 120  
 ctacgtgccc aggcctctct cctctcatc agatcacatg accttctctg agcgggtgaa 180  
 gaacatgctc attgcctttt cacagaactt tctgtgcgac gtggtttatt ccccgatgc 240  
 aacccttgcc tcagaattcc ttcagagaga ggtgactgtc caggacctat tgagctctgc 300  
 atctgtctgg ctgttttaga gtgactttgt gaaggattac cctaggcccc tcatgccccaa 360  
 tatggttttt gttggtggaa tcaactgcct tcacccaaat ccactatccc aggtgtgtat 420  
 tggagtggga cttttacatg cgtatattct ttcagatgta ttactttgga tcgattaact 480  
 agccccagat atatgctgag yaagcattct gagataattt aaaatgccct cttttgttaa 540  
 tttttgactc ctaggtttga gtctgtcttt ggcacatct tctggatgat ttcttggtat 600  
 ctgagatttc gggaaagcat tccttggaac ttttactctg tgtgctccag tggatagtaa 660  
 tcaattagaa acaacaagct gttaaagcc ataggcacag aatgctgggt ttggggcacc 720  
 ctgcagaaaa ctgagttgaa gcctgcacct tgccctggat tcagtcaggc aggcaatgtt 780  
 caggactgat gaaatcatc tttgatgat atagatcctg gaaatgaaag ttgcctttgt 840  
 gaccctggtt aaagctccag tttctaaata ttctgataag aagctaaatc ctgcagtcgg 900  
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<210> 293  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-405-54 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-405-54.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-405-54.mis2, potential complement

252

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<220>
<221> primer_bind
<222> 448..465
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 848..867
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-405-54 potential probe

<400> 293
tggtttttcc ctgccacttc ccaactatta atccaaagggt tttttttgtt gttgtggttg      60
ttgtcattgt tttcaatttg actctcaaat actctattaa actatgatcc accacactca      120
gaagtatcat tttctctaag agactcaaaa gtgtattagg gagaatttat ttaaaaaataa      180
aataaatggg atattgtttc ttcattattaa atagaagtat ttctccaaaa agctgttggt      240
tagaacactg aatttatgtc ttacatttct gctcttatag ttctgcatcc acttgtttca      300
ttaagcaaac tttcccttaa agtgcaggaa agtgaaaaaa tcctaagtgc acagcttgat      360
aaattatcac aaattcacgt agtgcataca cccttgtaac taaacctcca aaacaagatg      420
ccggaagttg ccagtcctca gaagccttca cagttactga tcctcccact ctgttaaaga      480
ctgttccttc agaggacccc ygttttctag ttagtatagc agatttggtt tctaatacata      540
ttatgttctt tctttacgtt ctgctctttt tgccccctcc aggtcctgtg gcggtacact      600
ggaacccgac catcgaatct tgcgaacaac acgatacttg ttaagtgggt accccaaaac      660
gatctgcttg gtatgttggg cggattggat gtataggtca aaccaggggc aaattaagaa      720
aatggcttaa gcacagctat tctaaaggat tggtgagctt gaaaatatta tggccaacat      780
atcctacatt gctttttatc tagtggggta tctcaaccca cattttcttc tgcaaatttc      840
tgcaagggca tgtgagtaac actgagtcct tggagtgttt tcagaaccta gatgtgtcca      900
gctgtgaaac tcagagatgt aactgctgac atcctcccta ttttgcatct caggtcaccc      960
gatgacccgt gcctttatca cccatgctgg ttcccatggg g                                1001

<210> 294
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-408-356 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 10-408-356.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-408-356.mis2, potential complement

<220>
<221> primer_bind
<222> 146..165
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 546..565
<223> downstream amplification primer, complement

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<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-408-356 potential probe

```

<400> 294
agaacatcat ggcctctccc agccttcaca aggaccgccc ggtggagccg ctggacctgg      60
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cccacgacct cacctggtag cagtaccatt ccttggacgt gattggtttc ctcttggccg      180
tcgtgctgac agtggccttc atcaccttta aatgttgtgc ttatggctac cggaaatgct      240
tggggaaaaa agggcgagtt aagaaagccc acaaatccaa gacctattga gaagtgggtg      300
ggaaataagg taaaattttg aaccattccc tagtcatttc caaacttgaa aacagaatca      360
gtgttaaatt cattttattc ttattaagga aatactttgc ataaattaat cagccccaga      420
gtgctttaaa aaattctctt aaataaaaaa aatagactcg ctagtcagta aagatatttg      480
aatatgtatc gtgccccctc ygggtgtctt gatcaggatg acatgtgccg tttttcagag      540
gacgtgcaga caggctggca ttctagatta cttttcttac tctgaaacat ggctgtttg      600
ggagtgcggg attcaaagggt ggtcccaccg ctgcccctac tgcaaattggc agttttaatc      660
ttatcttttg gcttctgcag atgggtgcaa ttgatcctta accaataatg gtcagtcctc      720
atctctgtcc tgcttcatag gtgccacctt gtgtgtttta agaagggaag ctttgtacct      780
ttagagtgtg ggtgaaatga atgaatggct tggagtgcac tgagaacagc atatgatttc      840
ttgctttggg gaaaaagaat gatgctatga aattgggtggg tgggtgtattt gagaagataa      900
tcattgctta tgtcaaatgg agctgaattt gataaaaacc caaataacag ctatgaagtg      960
ctgggcaagt ttactttttt tctgatgttt cctacaacta a                                1001

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<210> 295  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-409-148 : polymorphic base G or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-409-148.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-409-148.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 354..372  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 779..798  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-409-148 potential probe

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<400> 295
ctcacctggt accagtacca ttccctggac gtgattgggt tcctcttggc cgtcgtgctg      60
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254

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aaagggcgag ttaagaaagc ccacaaatcc aagacccatt gagaagtggg tgggaaataa 180
ggtaaaatTT tgaaccattc cctagtcatt tccaaacttg aaaacagaat cagtgttaaa 240
ttcattttat tcttattaag gaaatacttt gcataaatta atcagcccca gagtgcttta 300
aaaaattctc ttaataaaaa ataatagact cgctagtcag taaagatatt tgaatatgta 360
tcgtgcccc cccggtgtct ttgatcagga tgacatgtgc catttttcag aggacgtgca 420
gacaggctgg cattctagat tacttttctt actctgaaac atggcctgtt tgggagtgcg 480
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tggcttctgc agatggttgc aattgatcct taaccaataa tggtcagtcc tcatctctgt 600
cctgcttcat aggtgccacc ttgtgtgttt aaagaaggga agctttgtac ctttagagtg 660
taggtgaaat gaatgaatgg cttggagtgc actgagaaca gcatatgatt tcttgctttg 720
gggaaaaaga atgatgctat gaaattggtg ggtggtgtat ttgagaagat aatcattgct 780
tatgtcaaat ggagctgaat ttgataaaaa cccaaaatac agctatgaag tgctgggcaa 840
gtttactttt tttctgatgt ttcttacaac taaaaataaa ttaataaatt tatataaatt 900
ctatttaagt gttttcactg gtgtcgcatt tatttcttgt taagttgcat tttctaatta 960
caaaagtaat gcatgattat gacagaaagt ttggaaaata t 1001

```

&lt;210&gt; 296

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-409-249 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-409-249.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-409-249.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 253..271

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 678..697

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-409-249 potential probe

&lt;400&gt; 296

```

ccggaaatgc ttggggaaaa aagggcgagt taagaaagcc cacaaatcca agacccattg 60
agaagtgggt gggaaataag gtaaaatTTT gaaccattcc ctagtcattt ccaaacttga 120
aaacagaatc agtgtaaat tcatTTtatt cttattaagg aaatactttg cataaattaa 180
tcagccccag agtgctttaa aaaattctct taaataaaaa taatagactc gctagtcagt 240
aaagatattt gaatatgtat cgtgccccct ccggtgtctt tgatcaggat gacatgtgcc 300
atttttcaga ggacgtgcag acaggctggc attctagatt acttttctta ctctgaaaca 360
tggcctgttt gggagtgcgg gattcaaagg tgggtccacc gctgccccta ctgcaaatgg 420
cagttttaat cttatctttt ggcttctgca gatggttgca attgatcctt aaccaataat 480
ggtcagtcct catctctgtc stgcttcata ggtgccacct tgtgtgttta aagaagggaa 540
gctttgtacc tttagagtgt aggtgaaatg aatgaatggc ttggagtgcg ctgagaacag 600
catatgattt cttgcttttg ggaaaaagaa tgatgctatg aaattggtgg gtggtgtatt 660

```

255

tgagaagata	atcattgctt	atgtcaaagt	gagctgaatt	tgataaaaac	ccaaaataca	720
gctatgaagt	gctgggcaag	tttacttttt	ttctgatgtt	tcctacaact	aaaaataaat	780
taataaaattt	atataaaattc	tatttaagt	ttttcactgg	tgctgcattt	atctcttggt	840
aagttgcatt	ttctaattac	aaaagtaatg	catgattatg	acagaaagtt	tggaaaatat	900
agagggttcac	acacacatgc	cttcattgcg	tgtgcatgca	taaattgcatg	agaaaagaaa	960
aataaccagt	aatcgcatcg	cccagaaata	accccagtta	c		1001

&lt;210&gt; 297

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-410-274 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-410-274.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-410-274.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 228..245

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 645..664

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-410-274 potential probe

&lt;400&gt; 297

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accaataatg	gtcagtcctc	atctctgtcc	tgcttcatag	gtgccacctt	gtgtgtttaa	120
agaagggaag	ctttgtacct	ttagagtgtg	ggtgaaatga	atgaatggct	tggagtgcac	180
tgagaacagc	atatgatttc	ttgctttggg	gaaaaagaat	gatgctatga	aattggtggg	240
tggtgtattt	gagaagataa	tcattgctta	tgtaaatgg	agctgaattt	gataaaaacc	300
caaaatacag	ctatgaagt	ctgggcaagt	ttactttttt	tctgatgttt	cctacaacta	360
aaaataaatt	aataaattta	tataaattct	atttaagtgt	tttactgggt	gtcgcattta	420
tttcttggtt	agttgcattt	tctaattaca	aaagtaatgc	atgattatga	cagaaagttt	480
ggaaaatata	gaggttcaca	macacatgcc	ttcattgctg	gtgcatgcat	aaatgcatga	540
gaaaagaaaa	ataaccagta	atcgcatcgc	ccagaaataa	ccccagttac	aattgtggca	600
aatacacata	cttataaata	ttgcagatat	attaagtata	cctagtattt	gctaacactc	660
tttcttctac	tctgtcatga	agattctccc	aagggtgtttt	tgtataatat	ttaattcatt	720
ttcagtgccc	aagcagtatt	ctacttcatg	gatataccag	gatttattta	accataactt	780
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tcgggagttt	gagaccagcc	tggccaacat	ggcaaaaccc	cgtctctact	aaaaatacac	960
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&lt;210&gt; 298

&lt;211&gt; 1001

256

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 10-410-280 : polymorphic base C or T

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 10-410-280.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 10-410-280.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 222..239  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 639..658  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 10-410-280 potential probe

<400> 298  
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gaagctttgt accttagag ttaggtgaa atgaatgaat ggcttggagt gcactgagaa 180  
cagcatatga tttcttgctt tggggaaaaa gaatgatgct atgaaattgg tgggtggtgt 240  
atltgagaag ataatcattg cttatgtcaa atggagctga atltgataaa aacccaaaat 300  
acagctatga agtgctgggc aagtttactt ttttctgat gtttcctaca actaaaaata 360  
aattataaaa tttatataaa ttctatttaa gtgttttcac tgggtgctgca tttatttctt 420  
gttaagttgc attttctaata taaaaagta atgcatgatt atgacagaaa gtttggaaaa 480  
tatagaggtt cacacacaca ygccttcatt gcgtgtgcat gcataaatgc atgagaaaag 540  
aaaaataacc agtaatcgca tcgcccagaa ataaccccag ttacaattgt ggcaaataca 600  
catacttata aatattgcag atatatataag tatacctagt atltgctaac actctttctt 660  
ctactctgtc atgaagattc tcccaagggtg ttttgtata atatttaatt cattttcagt 720  
ggccaagcag tattctactt catggatata ccaggattta ttttaaccata acttctggtt 780  
ggattcactc ttattatttt gttaatttaa aaaaaaaaaga cctcggctgg gcacagtggc 840  
tcatgcctgt aatcccagca ctttgggagg ccgaggtggg tggatcacct aagatcggga 900  
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agccgggtgt ggttgccagc acctgtaatt ccagctaatt g 1001

<210> 299  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 10-410-337 : polymorphic base A or G

<220>

257

<221> misc\_binding  
 <222> 481..500  
 <223> 10-410-337.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-410-337.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 165..182  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 582..601  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-410-337 potential probe

<400> 299  
 aataatgggc agtcctcatc tctgtcctgc ttcataaggc ccaccttggt tgtttaaaga 60  
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 gaacagcata tgatttcttg ctttggggaa aaagaatgat gctatgaaat tgggtgggtgg 180  
 tgtatttgag aagataatca ttgcttatgt caaatggagc tgaatttgat aaaaacccaa 240  
 aatacagcta tgaagtgtcg ggcaagttta ctttttttct gatgtttcct acaactaaaa 300  
 ataaattaat aaatttatat aaattctatt taagtgtttt cactgggtgc gcatttattt 360  
 cttgttaagt tgcattttct aattacaaaa gtaatgcagc attatgacag aaagtttgga 420  
 aaatatagag gttcacacac acatgccttc attgcgtgtg catgcataaa tgcattgagaa 480  
 aagaaaaata accagtaatc rcacgcacca gaaataaccc cagttacaat tgtggcaaat 540  
 acacatactt ataaatattg cagatatatt aagtatacct agtatttgct aacactcttt 600  
 cttctactct gtcattgaaga ttctcccaag gtgtttttgt ataataattta attcattttc 660  
 agtggccaag cagtattcta cttcatggat ataccaggat ttatttaacc ataacttctg 720  
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 ggagtttgag accagcctgg ccaacatggc aaaaccccgct ctctactaaa aatacagaaa 900  
 attagccggg tgtggttgcc agcacctgta attccagcta attggggaggc tgaggcagga 960  
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<210> 300  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-121-326 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-121-326.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-121-326.mis2, complement

258

<220>  
 <221> primer\_bind  
 <222> 178..196  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 637..656  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-121-326 potential probe

<400> 300  
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 ccgcgcgaca ggtcagaggc ttggcgacct gggccgcctg gagggccgcc ctttatgacg 180  
 cagccacatc tcattggccg aggcctgtga gcgcctcgca tcccaagatg cagtgtcct 240  
 gggactggcc ctgctctctg tgaggctctg tgaggccctg tgatgctcca agaccaggcc 300  
 ccgccactc cggcctccaa ccagccatgg tctccaaaaa ggatgggaaa aagaggttgg 360  
 ggaaaagaga gggccttgac tttggctgcc tgaagaactg tttttcttaa agtaggcttt 420  
 atatcagtct ttttcctcgg ccacaggagg gaagagggtg gtgggagtga gtttagtctg 480  
 accggggctg aagacatcct rttgtttagg actgcggttc tccaacgttc cagccccggt 540  
 gccatttgc ttttgttcat ctggattatg cctatcatat gtactgcatt agagattaaa 600  
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 acaaaccttt ttatgtaact ttttttgaga caaaatgtag tgagaagagt ggcacgttt 720  
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<210> 301  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-122-341 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-122-341.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-122-341.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 162..180  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 595..612  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-122-341 potential probe

<400> 301  
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 gcagatcgga gctgttccta tttggccatt ggtgctgaag cttccgttct cctctccaac 180  
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 aggtcacact agatttaata tggtttggt cttgtgtccc caccaaatct catcttgaat 480  
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 aatctcatca tagtgagtga gttcttatga gatctgatgg ttttataagg ggcccttate 600  
 ccttcacttg caagctattc ctctttctct ctttcatgta agacgtgtct ttgcctctct 660  
 ctcactttct gccaggatta taagtttctt gaggcccctc cagccacatg gaactggagt 720  
 caattaaagc tctttccttt ataaattacc cagtcttggg tatgtcttta tagcagtgtg 780  
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 tgaaaatgtg gaagtgactt ggaactgggt aacaggcaga ggctggaaca gtttgagag 900  
 ctcaagagac aggaagatgt gggaaagttt ggaacttcct ggagacttgt tgaacggttt 960  
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<210> 302  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-122-381 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-122-381.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-122-381.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 122..140  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 555..572  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-122-381 potential probe

<400> 302  
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 cgtcgatcat gctgggagct gcagatcgga gctgttccta tttggccatt ggtgctgaag 120

260

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cttccgttct cctctccaac acctcctttt tttttccttt tctcttcag tttttcaatg 180
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cagaacaaat tgggaactgt ggcacagggt actcttgcta tgccccctaa gaattccaca 300
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cagccacatg gaactggagt caattaaagc tctttccttt ataaattacc cagtcttggg 720
tatgtcttta tagcagtgtg tgaatggact aatacagtaa attggtgtca tggagagtgg 780
ggtactgcta taaaactact tgaaaatgtg gaagtgactt ggaactgggt aacaggcaga 840
ggctggaaca gtttgagag ctcagaagac aggaagatgt gggaaagttt ggaacttcct 900
ggagacttgt tgaacggttt tgacaaaaat gctgataatg ttgtggacaa tgaagtccag 960
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&lt;210&gt; 303

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-124-169 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-124-169.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-124-169.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 334..352

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 830..848

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-124-169 potential probe

&lt;400&gt; 303

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gacatgttct ccttctgtca gagaagactt gggagttaca aagtctatag ctgtgtccat 180
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ctccctcaag aagaccggg kgtctagctg gagcctgcgc ttgagcaaag agctgaacaa 540
ccagattgag agctttgaca gcccctctct ggagaagggt tgtcttcata gggacaggag 600
agtttgagca aggagtagac tccaggggtg atggagtgga gaggataaca ccgtgtgtgt 660

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261

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gagatcagct ggacctgtgg ctgtttacag ggtcctggga ggagaggagc gtgggtctgga 720
ttgggtgggtt tagttttgtt ttgggtttatt tttttgcttt gttttgttga tcctccagat 780
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caaaattcta aaacattttt gaaagaaatc agtgagggtc tcaataaata gaaagcacc 900
catgttcagtg gattgaaaga tgctatagtt caaatggtga tactctccaa atttatcttc 960
agtttttagca tgacacctat cacaatccaa gctagttctt t 1001

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&lt;210&gt; 304

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-124-194 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 12-124-194.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-124-194.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 309..327

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 805..823

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-124-194 potential probe

&lt;400&gt; 304

```

caagtttgct gaggcctggg aggaggagaa agatgggagc agatggaaaag gcatttggtg 60
ggctgggcac atgaggagtt tagaagcccc taaaggacat gttctccttc tgtcagagaa 120
gacttgggag ttacaaaagtc tatagctgtg tccatcccca ggggagaact gagagaacct 180
tccatccccga gagccaggtg gccaggaggg cagcaggagg tgacaccctt tctgctagcc 240
gccagagcca cagccattcc tcagaccacac agaacattat tagcactgga tcagggatct 300
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aggtcacgta gcattcccag agcccaggca agaacaagga tgaaggcctt tggcccttct 420
caccaacgtc ttgtgcctcc ccagtttctg cgaaactccc tcaagaagac ccgggggtct 480
agctggagcc tgcgcttgag yaaagagctg aacaaccaga ttgcgagctt tgacagcccc 540
tctctggaga aggtttgtct tcatagggag aggagagttt ggacaaggag tagactccag 600
gggtgatgga gtggagagga taacaccgtg tgtgtgagat cagctggacc tgtggctggt 660
tacaggggcc tgggaggaga ggagcgtggg ctggattggt gggttttagt ttgttttggg 720
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tccaaagctag ttcttttttt tgatcctaaa attgatccta a 1001

```

&lt;210&gt; 305

&lt;211&gt; 1001

262

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-124-300 : polymorphic base G or T

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-124-300.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-124-300.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 203..221  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 699..717  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-124-300 potential probe

<400> 305  
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 cccttctgct agccgccaga gccacagcca ttctcagac ccacagaaca ttattagcac 180  
 tggatcaggg atctatggac ttagtgatct ggagatgtgg gcaaggggag cgagagaggg 240  
 caagtcagta ctcaaggtca cgtagcattc ccagagccca ggcaagaaca aggatgaagg 300  
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 gctttgacag cccctctctg gagaagggtt gtcttcatac ggacaggaga gtttggacaa 480  
 ggagtagact ccaggggtga kggagtggag aggataacac cgtgtgtgtg agatcagctg 540  
 gacctgtggc tgtttacagg gtcttgggag gagaggagcg tggctctggat tgggtggttt 600  
 agttttgttt tgggtttattt ttttgccttg ttttgttgat cctccagatt ctggctagca 660  
 atcacatgtg tgtatttctt tgtaaacaca aaactatacc gcattgctac aaaattctaa 720  
 aacatttttg aaagaaatca gtgaggggtct caataaatag aaagcaccac atgttcatgg 780  
 attgaaagat gctatagttc aaatgggtgat actctccaaa tttatcttca gttttagcat 840  
 gacacctatc acaatccaag ctagtctctt tttttgatcc taaaattgat cctaaaattt 900  
 atatggaaat accaggggct ctgaatagcc aacaaaatta tgagaagaag aataaagttg 960  
 gaggactcaa acttctcaat ttcaacttct tacacaactg c 1001

<210> 306  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-124-58 : polymorphic base A or C

<220>

263

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<221> misc_binding
<222> 481..500
<223> 12-124-58.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-124-58.mis2, potential complement

<220>
<221> primer_bind
<222> 444..462
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 940..958
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-124-58 potential probe

<400> 306
gccttgaata ccaggcttag gggagcagtg gaagaaagtg gcatgtgtga ccatacatgt      60
gttcctgcag aacataactga gttcacttgg gatcagaaaag cctgggaaga caagctgatt      120
caggtaaacg ggtaacaagt ttgctgaggg ctgggaggag gagaaagatg ggagcagatg      180
gaaaggcatt tgggtgggctg ggcacatgag gagtttagaa gcccctaaag gacatgttct      240
ccttctgtca gagaagactt gggagttaca aagtctatag ctgtgtccat cccaggggga      300
gaactgagag aaccttccat cccgagagcc aggtggccag gagggcagca gggagtgaca      360
cccttcttgc tagccgccag agccacagcc attcctcaga cccacagaac attattagca      420
ctggatcagg gatctatgga cttagtgatc tggagatgtg ggcaaggga gcgagagagg      480
gcaagtcagt actcaaggtc mcgtagcatt cccagagccc aggcaagaac aaggatgaag      540
gcctttggcc cttctacca acgtcttgtg cctccccagt ttctgcgaaa ctccctcaag      600
aagaccgggg ggtctagctg gagcctgcgc ttgagcaaag agctgaacaa ccagattgcg      660
agctttgaca gcccctctct ggagaagggt tgtcttcata gggacaggag agtttggaca      720
aggagtagac tccaggggtg atggagtgga gaggataaca ccgtgtgtgt gagatcagct      780
ggacctgtgg ctgtttacag ggtcctggga ggagaggagc gtggtctgga ttggtggttt      840
tagttttgtt ttggtttatt tttttgcttt gttttgttga tcctccagat tctggctagc      900
aatcacatgt gtgtatttct ttgtaaacac aaaactatac cgcattgcta caaaattcta      960
aaacattttt gaaagaaatc agtgagggtc tcaataaata g                                1001

<210> 307
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-126-222 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-126-222.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-126-222.mis2, potential complement

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264

<220>  
 <221> primer\_bind  
 <222> 703..722  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 267..286  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-126-222 potential probe

<400> 307  
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 ttattgtgat gacatgaccc tgtttatcta tggtaatgtc atttgctcta aaatgaactt 120  
 ttatatcaga acggcctctg gggccatctg gagtgcaggg tgggctgtgc tcttggccct 180  
 tcattacagg cagacatttg gagccaggcc aaatgcaaca gccaagcacc atcctgggtga 240  
 gaatgctggg gatgctcttt ccttgcctac agcctcatct cccttcatga tggggcctgt 300  
 ctagaaccac aaggaagcat tgaaagtcga tctgagtagg agggtgaggt gggcaaaagc 360  
 tggccaggaa agggctggcc ccaggatttt aacacccact ttggtgttct gggctctacc 420  
 catagctcct ggaaaagtgg atcttgtcgg agaaagaatg ggagcgggaa aaggccgtga 480  
 gcctccatct ctatctcatg yggatttatg tccacagcac tgctgtctgt gtgagtcag 540  
 gagtccccag cccccagcct gctgctctga catgcacaga aaaattcctg tgtctatctg 600  
 gaagatgagg gaacacttat gaaatcaagt tctgtcccca gagggcccct tgggagaggc 660  
 tgcagtgacc tagtgccctg ggctgggtca gggagggaga ggggtgggatt gtggttggca 720  
 gagctccagg ctttgtgggg cctgtggcct ctgtagcggt gagagccctt ttaagaaaa 780  
 aagatacata gttatgaacc ccacattggc taggaaaatg aatatttact ttgagaaaat 840  
 caagtgtgc aaaatcataa attttgcaaa gcttgacaag taccgcaacc cttaaaatgt 900  
 ctagaacgac gtagtgtttt tactgattga ccacttgaca cagcctgatg agacttaatt 960  
 gtctacactt tttggctcca tgctgtttga ctgtcttgtc c 1001

<210> 308  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-126-297 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-126-297.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-126-297.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 778..797  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 342..361  
 <223> downstream amplification primer

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<220>
<221> misc_binding
<222> 489..513
<223> 12-126-297 potential probe

<400> 308
gaaatctctg actaatagtg tgaacttggt tatttctcct tttggctctt tcattttttg      60
cttcatgtat tatttagcta tggtattagg tgcataaaca tttaggattg ttatgtctgg      120
ttgatgaact gacccttatt gtgatgacat gaccctgttt atctatggta atgtcatttg      180
ctctaaaatg aacttttata tcagaacggc ctctggggcc atctggagtg caggggtgggc      240
tgtgtctctg gcccttcatt acaggcagac atttggagcc aggccaaatg caacagccaa      300
gcaccatcct ggtgagaatg ctggggatgc tctttccctg cctacagcct catctccctt      360
catgatgggg cctgtctaga accacaagga agcattgaaa gtcgatctga gtaggagggg      420
gaggtgggca aaagctggcc aggaaagggc tggccccagg attttaacac ccactttggt      480
gttctgggct ctacccatag ytcttgaaa agtggatctt gtcggagaaa gaatgggagc      540
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tctgtgtgag tccaggagtc cccagccccc agcctgctgc tctgacatgc acagaaaaat      660
tcctgtgtct atctggaaga tgaggggaaca cttatgaaat caagttctgt cccagagggg      720
ccccttggga gaggctgcag tgacctagtg ccctgggctg ggtcagggag ggagaggggtg      780
ggattgtggg tggcagagct ccaggctttg tggggcctgt ggcttctgta gcggtgagag      840
ccctttttta gaaaaaagat acatagttat gaaccccaca ttggctagga aaatgaatat      900
ttactttgag aaaatcaagt gctgcaaaat cataaatttt gcaaagcttg acaagtaccg      960
caacccttaa aatgtctaga acgacgtagt gtttttactg a                                     1001

<210> 309
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-128-225 : polymorphic base T or G

<220>
<221> misc_binding
<222> 502..520
<223> 12-128-225.mis1, complement

<220>
<221> misc_binding
<222> 481..500
<223> 12-128-225.mis2, potential

<220>
<221> primer_bind
<222> 706..725
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 276..295
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-128-225 potential probe

<400> 309
ccctgaaatg ggaccatgac agctgggtct gagagacagt ggtagaaaaca tccagattca      60
gcatttactt gctggcttgg atgcaggggtc tagaacgaaa agagaagaaa agtcacttct      120

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266

atacagaaac	atgtccagag	cgcttactgt	ctccaaaacc	atggactggc	acctgagtga	180
tagcatgatt	ccaaagccaa	aatcttgcct	gtaaggaata	tatatatata	tatatatata	240
tatatatgta	tatatgatat	agctatagtc	taatagcaag	gacagatatg	caaactgcta	300
aaagatacaa	ggcagaacag	aacaaaatgc	tgtttttctg	ggatttttga	aattcaagga	360
attcaaggga	ttcaaggaag	gtggccttgc	ttcccgggag	ggtcctgtag	atgatctaca	420
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ttaaaaagaa	agtagcttca	aaagggttcc	agaaacactt	tccatggacg	tgtcactctt	600
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agagcaccat	cccctgtaat	tgcctgggtc	tgagtttgtc	tctgtctacc	tgacccctcc	720
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ccatccagta	gggctcacac	gttccataaa	tatttggcag	atgagggaat	tagcaatggg	840
ttctgctttg	gtttcagagc	agatattaat	tggattgctt	agtagtggtt	ctctgttgta	900
attcatgagc	atgaatgtgg	attgcccact	attcagatta	gtaagtattt	cttgggtcaag	960
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&lt;210&gt; 310

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-129-176 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-129-176.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-129-176.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 326..344

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 779..797

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-129-176 potential probe

&lt;400&gt; 310

ggttgatgcat	gtccttcatt	gggaaggagc	agggacacta	cattgagact	tgacccatct	60
ggattttttt	tttgcacaat	tttcagggga	aagatgatgc	aacagcaaat	tacaattgtt	120
aatgtgataa	tttttagtgg	tcctgtcttg	cgaatgatag	agaggtgacc	acaggagacc	180
taagcactcg	caggaagtag	aagtgttaaa	gaggttgact	cagttcagtg	gaagtagagc	240
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agggatctgt	gctgacaaaa	gatttttctg	gtcaggattt	ggggtctggg	gcatgatgtg	360
gggacatctc	agagttcggg	agggcatagt	ttggttgacg	gtcaattggg	tcttctactt	420
ggaatgctga	aattatcaag	aaattgtgga	aggggtctagg	gaggagataa	aactgtgagc	480
gtataagccc	agttaagctg	kggacgggtg	tgaatggaca	tgtgtccaag	aagggaagtg	540
tttctcaggt	gaagctgaac	atatcaccaa	acccacccta	tcccactcca	agtttctata	600
gtgggatcta	cttctttacc	aacaatttca	aaggcagctt	tctttaatcc	agaatatttg	660

267

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gggttcattg atgtggtact ctgggacctg aatattgttt cttattcctt ggtgtgccat 720
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atgacttgat ccttttgtat ttttatttct gtacttcctt ttattaacat aggtattatt 840
gccaaacact ctaagcttca ttttttaaaa tcaaaccaca tgattttttt attggtgatt 900
ttctccctta tgcagtgtag ttattcaatt ataaaatatg tgtttattag atttgtaaca 960
atgttgaaaa gtgggatgga atttaaata tagactcttgc c 1001

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&lt;210&gt; 311

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-130-203 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-130-203.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-130-203.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 301..319

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 733..753

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-130-203 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 167

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 311

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aatcgttctt tgaaacagaa catcttctga tgaaattttc tagaagaatg gcaattatga 60
acaatatgtc tttgatcata cataggtctt gtgtggagct actgcataat gaggccctga 120
tcaggcacct gcatgctact tcctttgatg tggttctaac agaccnctt tcacctctgc 180
gcggcggtgc tggctaagta cctgtcgatt cctgctgtgt ttttcttgag gaacattcca 240
tgtgatttag actttaaggc cacacagtgt ccaaaccctt cctcctatat tcctagatta 300
ctaacgacca attcagacca catgacattc ctgcaaaggc tcaagaacat gctctaccct 360
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ctttttcaga gagaggtgtc agtgggtgat cttgtcagcc atgcatctgt gtggctgttc 480
cgaggggact ttgtgatgga ytaccccagg ccgatcatgc ccaacatggt cttcattggg 540
ggcatcaact gtgccaacgg gaagccacta tctcaggtct gtattggtgc ctttatccaa 600
tcaatgttcc aggcaaaaca ctttttaaaa aatgtattta cttacaagtg cttccatata 660
tacttatctt tccaaagatt tcatttctgc ttctcattgt tgtaaatagtc ttcagtgaga 720
taaactttta aagggtcaat ggtagtgcag ttcagggttt caatggccac tgagagggaag 780
gagaggcagg gacgaggatc tgtcaaagga tgggcaagag tgggtgtgact cacggagact 840
gttcggttgc aaaagcacca tcttcatggc tgtggatgtg cttcagctgg gcaggagcag 900

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268

ggacactaca ctgagaactg atccatccaa tcttgctggc aagattttca gtagaaagga 960  
 ggaaggaata gaaatttaca attgttgacg tgacaatttt t 1001

<210> 312  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-130-260 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-130-260.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-130-260.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 244..262  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 676..696  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-130-260 potential probe

<220>  
 <221> misc\_feature  
 <222> 110  
 <223> n=a, g, c or t

<400> 312  
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 cctctggccc tgctctacct ttgccatgct gttctgtctc cttatgcaag ccttgccctc 360  
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 agataaactt ttaaagggtc aatggtagtg cagttcaggg tttcaatggc cactgagagg 720  
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 ggaggaagga atagaaattt acaattgttg acgtgacaat ttttagtggt cccatcttgc 960  
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<210> 313



269

<211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-131-112 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-131-112.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-131-112.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 594..612  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 104..124  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-131-112 potential probe

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 ctccccactc cctgttttaa gaaagctggc ttagcaatgt tgtctgcatt ttggatgtgc 180  
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 cagagaccat atggttgggg actagggcaa tgggtgactcc tcagacctca gctgcagcct 300  
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 gactccaggt tcccctgccg cagctggcca caggactgct gcttctcctc agtgtccagc 720  
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 ggaccagga cgaatttgat cgccttttgc tgggtcacac tcaatcgttc tttgaaacag 960  
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<210> 314  
 <211> 755  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 255  
 <223> 12-132-157 : polymorphic base C or T

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<220>
<221> misc_binding
<222> 235..254
<223> 12-132-157.mis1, potential

<220>
<221> misc_binding
<222> 256..275
<223> 12-132-157.mis2, potential complement

<220>
<221> primer_bind
<222> 99..117
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 557..577
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 243..267
<223> 12-132-157 potential probe

<400> 314
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cccctctcca cattcctccc accctcattc ccccttctcc tggaaacatc caagtccaaa      180
ttaatactat taatgcctac ctggatatta atgcctacct gatcatatct aggtcatggt      240
agaattgtgt aaacytcctt tggatagctc actcctcatg tcctgtcccc ccagcccaca      300
gccctgggtcc agatcccac cttgggagct cctggcccag cctttcccat gcatgacctt      360
actgggcctc tgctctgcct ggcttctctg agccagctag agtaactata ggcattcccat      420
cctctgtaga ctaccctcac tgtggtcac tcagagtggg cactgttaac accaggctct      480
caaaacaact atgcccacct accatgcttg ctctgtccc cttccctcca actctggccc      540
caagggcatc gctgggtgat taggtgattg atggggctct ttaggatcca gggctagcaa      600
agggcagagc ctgctctctg actttgatct aaacaagctg gccagtcaat cctctgaggt      660
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gcatcttctc agcaggcctg gccgtgccct tgccc                                755

<210> 315
<211> 991
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-132-437 : polymorphic base A or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-132-437.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-132-437.mis2, potential complement

<220>
<221> primer_bind
<222> 68..86

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271

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 526..546

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-132-437 potential probe

&lt;400&gt; 315

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cccttctcct	ggaacatcc	aagtccaaat	taatactatt	aatgcctacc	tggatattaa	180
tgcctacctg	atcatatcta	ggatcatgta	gaattgtgta	aacctccttt	ggatagctca	240
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ctggcccagc	ctttcccatg	catgacctta	ctgggcctct	gctctgcctg	gcttctctga	360
gccagctaga	gtaactatag	gcatcccatc	ctctgtagac	tacctcact	gtgggtccact	420
cagagtgggc	actgttaaca	ccaggctctc	aaaacaacta	tgccaccta	ccatgcttgc	480
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gttgagtcc	ctccaggtcc	cctgcactgg	catcttctca	gcaggcctgg	ccgtgccctt	720
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gcgcattggc	tgaaaagagc	atcttctaaa	cctcaaccca	ttagcacgat	tctccctctc	960
tcccaggata	agtgaatgt	aatgatgttc	t			991

&lt;210&gt; 316

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 666

&lt;223&gt; 12-133-153 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 646..665

&lt;223&gt; 12-133-153.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 667..686

&lt;223&gt; 12-133-153.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 800..818

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 250..270

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

272

&lt;222&gt; 654..678

&lt;223&gt; 12-133-153 potential probe

&lt;400&gt; 316

```

ggctctgggat ctaacgggtga tagttctagt gaaaccctcc cacccttcca ggagacccat    60
tcctataaga aatatgactg aagagcccag ctgtctcact ttgataacca attcttccat    120
ttgcatttcc attgatgtgt attgggccaa gaggagagct ttttcttggt gttaatatct    180
gtttaatcca gaagaagacc ctaaggaaag gtcagtattt aggtcccaga gtagctcttt    240
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ggatcaagga acttgggaca ggaggagaaa ggatctgaga gcctttcttc caccatctcg    360
acatctatgt tgggtgttct caatttttaa gaagctgtgg atgttgagag aacccttaa    420
aggagtattg agagatactg atatgaattt ctcattagcc attaattatc agtgtgtaga    480
agagagggcg agagggtctt attaaactaa ggatattatc tattgctgtg taacaaatta    540
tcataaaactc agtggcttaa aagaacatcc attgattatc tcacgggtcc actggccagg    600
agtctaggca tggctttcct tggctcttg tttaggggtc cagagactgc agtcctttct    660
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gtaggactga ggccctcagt tccttgctac gtggccctca ccacaggcag tttgcatcat    780
ggctgtttac ttttcaaggc cagtagaatg tttctgttca atctgataag agggaatctc    840
atgtggcata gtgtgatcat gggaatgaca tcccaccatc tttgcatat gatgtaatcc    900
actcgagaga gggtcaccca tcacctttgc catattctgt tgattagaag gaagtcacag    960
gttccaccca cacaccctca aggggatggc tttatggaag g                                     1001

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&lt;210&gt; 317

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-133-318 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-133-318.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-133-318.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 800..818

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 250..270

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-133-318 potential probe

&lt;400&gt; 317

```

ggctctgggat ctaacgggtga tagttctagt gaaaccctcc cacccttcca ggagacccat    60
tcctataaga aatatgactg aagagcccag ctgtctcact ttgataacca attcttccat    120
ttgcatttcc attgatgtgt attgggccaa gaggagagct ttttcttggt gttaatatct    180
gtttaatcca gaagaagacc ctaaggaaag gtcagtattt aggtcccaga gtagctcttt    240
aaaatccttt aaaggaaaag agagtatggg atagtaaagg gccatgggat ggatctttga    300

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273

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ggatcaagga acttggggaca ggaggagaaa ggatctgaga gcctttcttc caccatctcg 360
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aggagtatat agagatactg atatgaattt ctcattagcc attaattatc agtgtgtaga 480
agagagggcg agagggctct wttaaactaa ggatattatc tattgctgtg taacaaatta 540
tcataaactc agtggcttaa aagaacatcc attgattatc tcacgggtcc actggccagg 600
agtctaggca tggctttcct tggctctttg tttagggctc cagagactgc agtcctttct 660
ggagttcagc ttcctcttcc aagctcatgt gcttattgga gaattcagtt tcttatggtt 720
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actcgagaga gggtcaccca tcacctttgc catattctgt tgattagaag gaagtcacag 960
gttcaccca cacacctca aggggatggc tttatggaag g 1001

```

&lt;210&gt; 318

&lt;211&gt; 643

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 249

&lt;223&gt; 12-136-238 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 229..248

&lt;223&gt; 12-136-238.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 250..269

&lt;223&gt; 12-136-238.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 12..32

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 442..461

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 237..261

&lt;223&gt; 12-136-238 potential probe

&lt;400&gt; 318

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acacaggggtg agcaaaaggg agacttttta tccttttttg agaaagcaac cacaaaattt 60
tgaccttatg aagtatcttt gaaactttcc aaagctgatt agaatagcgt tgatgtattt 120
taatcagata cagaaagact ttacaaagta tgagctgcac acttgctgtc gtaatgctta 180
taagcactaa tttttttgta tacattcatg tacttgtaat tgcacaaaat gcgaaggaca 240
catttggcrg tgaggatgct gtgcacaatc aactccta atttttcagg aaaggataaa 300
cattccgcta aaaaatgagga aagacaagaa atgttcaaaa gcatgtcaaa agtgtccagc 360
tgttttcata gaatataggt gatttccttg tgctagcaaa atccagaaga aacgttataa 420
taactagaaa tgtctataat tgtgaggaag cccaagtatg aagattcaca aaaatagaat 480
aagtggatga aattagttaa aacaaccttc ttagtaatga ggtactctga ttataaaagt 540
actttgactc tttagaatgt gtcactgaac accaagctga aaaaacaaat taacagtctg 600
tcactagagc atgatctccc acagacaata tgctcataaa tga 643

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&lt;210&gt; 319

&lt;211&gt; 1001

274

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-138-141 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-138-141.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-138-141.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 623..641  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 186..204  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-138-141 potential probe

<400> 319  
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gttttgggagc cattgtccta cagcacatga gctgagttgc aaagatgttg agccagtggg 180  
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gcctagctgg gttaaggggc tggggtagag gatgttggag ggagtgatac aaagtgaac 300  
caggggacct ggtgggcttg tacagacaag gtttgcatac cgtgggtacca gacaaggttt 360  
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gacacataac aattgtacat atttatagaa tacaaagtga tatttcaaaa catatataca 720  
atgtgtaatg accgaatcag ggtaattagt atatccaaca ctgcaaacad ttatcatttc 780  
tttgtgttgc aagcattcaa aactgtttct tctagttttt tgaaaatata caatcaatca 840  
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<210> 320  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-138-42 : polymorphic base T or G

<220>

275

<221> misc\_binding  
 <222> 481..500  
 <223> 12-138-42.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-138-42.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 524..542  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 87..105  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-138-42 potential probe

<400> 320  
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 gggagtgata caaagtgcaa ccaggggacc tgggtgggctt gtacagacaa ggtttgcattc 240  
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 ttcccccttt tatttttagt tgacacataa caattgtaca tatttataga atacaaagtg 600  
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 actgcaaaca tttatcattt ctttgtgttg caagcattca aaactgtttc ttctagtttt 720  
 ttgaaaatat acaatcaatc attgttgact atattcacaa cctacagtgc tatagaacac 780  
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<210> 321  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-138-67 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-138-67.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-138-67.mis2, potential complement

276

<220>  
 <221> primer\_bind  
 <222> 549..567  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 112..130  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-138-67 potential probe

<400> 321  
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 gatacaccag acacagtgtg cagggatgtt tggctagtgg ggagatgcct agctgggtta 180  
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 aagtctgggg tggagcctga gaatcagcat ttctaacaag tcccaggtg atgccaaggc 420  
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 gggtagcaaa ggggtgctat ragcagagga gcttatttat aaaaagtttt attatttttg 540  
 ttggatttgg atttgcattt gcgttttccc ccttttattt ttagttgaca cataacaatt 600  
 gtacatattt atagaataca aagtgatatt tcaaaacata tatacaatgt gtaatgaccg 660  
 aatcagggtg attagtatat ccaacactgc aaacatttat catttctttg tgttgcaagc 720  
 attcaaaact gtttcttcta gttttttgaa aatatacaat caatcattgt tgactatatt 780  
 cacaacctac agtgctatag aacacaagag cttattcctt ctctccagct gtaattttgt 840  
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<210> 322  
 <211> 663  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-139-380 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-139-380.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-139-380.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 122..139  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 602..620  
 <223> downstream amplification primer, complement



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<220>
<221> misc_binding
<222> 489..513
<223> 12-139-380 potential probe

<220>
<221> misc_feature
<222> 655,658
<223> n=a, g, c or t

<400> 322
caagtctgac acctgtgtgg caggccagca ggctggagac tagcaggggc ggatgctgca      60
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tcacccacct atgttattga ggataatctc ttttatggat tatgcacttt cgtcacacct      180
acaagctacc ttcacaacaa cacctagatt ggtgtttgac tgaatgacta ggtacagtcg      240
cctagccaag ttgacacaca aaactgaccg tcataccagg catgcacaac tggatcatct      300
gccccggggtg cccctctaga ggacttggga tcagagggag ggccctggat gagaaggggt      360
tgatgagtgt gggtttgtat gaggctggca gtgataggag ttgggcaaca gggatgctgg      420
ccccagcagg gactagcaga ctctcttgcc acatcagggt gggtgacact taaatgcccc      480
tgacaaagac tggatttggc rgtggagcca gttcagggt gagactgtca ggggacatct      540
ctggggaagc agggatggga tggcaccatg gcagagcctg actgttctcc ttagaagccc      600
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agg                                                                                   663

<210> 323
<211> 951
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-140-134 : polymorphic base G or T

<220>
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<222> 481..500
<223> 12-140-134.mis1, potential

<220>
<221> misc_binding
<222> 502..520
<223> 12-140-134.mis2, complement

<220>
<221> primer_bind
<222> 368..386
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 868..888
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-140-134 potential probe

<220>
<221> misc_feature
<222> 923

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<223> n=a, g, c or t

<400> 323

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ctgtagatca ttctggctaa ataatgggat tcacttttaa gtgaaggagg ccttgaatgc      60
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ggctggggtga aatgcaggct tggattcagc aggaccgggg tagggcctga gagtctgcat      180
ttctaatacag ctcccaggga atgctgggtgc tcagtgggtc agtgggcccc tgagtagcga      240
gggtctagag tttgaccttg agttggtaga aggtcacctg ccaccatttg ccaatccact      300
gcagggtcttg aggcagatcc tggaaactgtc agtcaccacc aacacccttg tccccaaat      360
gcagctacac accattttca cagaactgca cgtccagggtg aggccagcaa gctcagctgg      420
acaagggcat tctctgggtg gatcagcagg aagctggcgg cgggaggtgg cctggcccac      480
aactcactcg cgtgcttcct ktatcctgat ggggtatgag gctgtgggtgc ccatgcagcc      540
ccagggtatgg tcagctgggc tgcccctgca cctgtgtccc tgtgatctcc tccagcctca      600
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gcctgggact gccttgcctg gcctgctctg gaccgcgtga gcatgactaa agctgtcctt      840
ccaccctgca gctcgtcctt accccaagga gctgatgaag ttcttcttca gccagatgga      900
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```

<210> 324

<211> 756

<212> DNA

<213> Homo Sapiens

<220>

<221> allele

<222> 501

<223> 12-140-329 : polymorphic base C or T

<220>

<221> misc\_binding

<222> 481..500

<223> 12-140-329.mis1, potential

<220>

<221> misc\_binding

<222> 502..521

<223> 12-140-329.mis2, potential complement

<220>

<221> primer\_bind

<222> 173..191

<223> upstream amplification primer

<220>

<221> primer\_bind

<222> 673..693

<223> downstream amplification primer, complement

<220>

<221> misc\_binding

<222> 489..513

<223> 12-140-329 potential probe

<220>

<221> misc\_feature

<222> 728

<223> n=a, g, c or t

<400> 324

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agggaatgct ggtgctcagt gggtcagtgg gccctgagt agcgagggtc tagagtttga      60
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279

gatcctggaa	ctgtcagtca	ccaccaaacac	ccctgtcccc	caaatgcagc	tacacacccat	180
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gggtggatca	gcaggaagct	ggcggcggga	ggtggcctgg	cccacaactc	actcgcgtgc	300
ttcctttatc	ctgatggggt	atgaggctgt	ggtgcccattg	cagccccagg	tatggtcagc	360
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caggtgtgca	acaaggcccc	ggcccagcat	cagtacagca	gccagaatct	gatggagatg	480
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ggcctgcagg	aggcctctag	gagtcccggc	ccctctgagc	gctgggcctg	ggactgcctt	600
gctgggcctg	ctctggaccc	ggtgagcatg	actaaagctg	tccttccacc	ctgcagctcg	660
ctcctacccc	aaggagctga	tgaagttcct	cttcagccag	atggagacaa	acaaggaggc	720
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&lt;210&gt; 325

&lt;211&gt; 700

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-140-385 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-140-385.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-140-385.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 117..135

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 617..637

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-140-385 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 672

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 325

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ccattttcac	agaactgcac	gtccaggtga	ggccagcaag	ctcagctgga	caagggcatt	180
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gtgcttcctt	tatcctgatg	gggtatgagg	ctgtgggtgcc	catgcagccc	caggtatggt	300
cagctgggct	gccccgtcac	ctgtgtccct	gtgatctcct	ccagcctcac	tgcacgtccc	360
ttaccaggtg	tgcaacaagg	ccccggccca	gcacagtagc	agcagccaga	atctgatgga	420
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ccgaggcctg	caggaggcct	staggagtec	cggccctct	gagcgctggg	cctgggactg	540
ccttgctggg	cctgctctgg	acccggtgag	catgactaaa	gctgtccttc	caccctgcag	600

280

ctcgctccta cccaaggag ctgatgaagt tcttcttcag ccagatggag acaaacaagg 660  
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<210> 326  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-141-159 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-141-159.mis1, potential complement

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-141-159.mis2

<220>  
 <221> primer\_bind  
 <222> 640..658  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 181..200  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-141-159 potential probe

<400> 326  
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 cgggtcatcaa ggcagaaccg actgacaacc tgggttctcc agtgcgagcc ttggcgatgg 120  
 aggccctctc gcacctgagg tgagctgggt tcccaccctc accccatccc aaggggtagg 180  
 gaaaagtcca agaccattcc tcggtgctcc ttcaggggtt gtgcctcctt tttcctctcc 240  
 tccctctgta gcaggggtct cacttggcgg ctccccagg caatgggcca gcctagatga 300  
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 tctccttga tttatggcac atgggtgggc tgctcttcca ttgattgggg atattttgag 420  
 caaaataaca gtgataacac tgaagagcat tgcaagcatc cttcaagatg agtcaggctc 480  
 agtgagtgtg ttgaagggtc yagtcaacta tttgttgata ccaactggtac aaaatgatct 540  
 aaagcccaag acttaccttc ctggagatgc caatccattg gtggggaggg ggaaagggga 600  
 agaagttatg gatttataaa attagtattt aaagagtttg ggtcatctct aagaatacct 660  
 cccctatagg ggaagattca ttacttaggc gcatagcaga actatctgtt actttaagct 720  
 gcatcaaaaa ggcagatttt aaaataaatg tcatgaacat caactaagca caatgatatg 780  
 ctgagtactg aaggacaca gagatacata agaccatgac ccagcctttt aaggacttag 840  
 agataagaca gacacataca aacaggatta taggtcaaag ctgactaaca ggactgtgtt 900  
 tacaggtaaa cccacacaca catcagcaaa aaaatatctg agaacaaaaa aaaaggggct 960  
 actaacaaaa gcaactcagt ttatacaaaa ggagctatca a 1001

<210> 327  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>

281

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<221> allele
<222> 501
<223> 12-141-392 : polymorphic base C or A

<220>
<221> misc_binding
<222> 481..500
<223> 12-141-392.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-141-392.mis2, potential complement

<220>
<221> primer_bind
<222> 873..891
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 414..433
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-141-392 potential probe

<400> 327
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acacagcaag tagatgagca gcgcgagctg ggagcccagg ccacttttgg gccctgctcc      120
cactctgctg cctcccctaa gatgccaaaa gccaaatggc ctaaaaaaga atgaaagaga      180
tgtgggggtg tagacacttt gggaagtgcc aggtcttcct catgctaaca ctggggagtg      240
ctgccgccat catggtcacc aaaagccacc acccctgct tgtgttgga aggcgggcat      300
caaggcagaa ccgactgaca acctggtttc tccagtgcga gccttggcga tggaggccct      360
ctcgcacctg aggtgagctg gggtcccacc ctcaccccat cccaaggggt agggaaaagt      420
tcaagaccat tcctcgggtc tccttcaggg gttgtgcctc ctttttcctc tcctccctct      480
gtagcagggg tctcacttgg mggctcccca gggcaatggg ccagcctaga tgatgggggt      540
cggcctcagc cactcttttc tataattctc agccagattt caagttgggt gcttctcctt      600
ggatttatgg cacatgggtg ggctgctctt ccattgattg gggatatatt gagcaaaata      660
acagtataaa cactgaagag cattgcaagc atccttcaag atgagtcagg ctcaagtgagt      720
gagttgaagg gctcagtcaa ctatttgttg ataccactgg taaaaaatga tctaaagccc      780
aagacttacc ttcctggaga tgccaatcca ttgggtggga gggggaaaagg ggaagaagtt      840
atggatttat aaaattagta tttaaagagt ttgggtcatc tctaagaata cctcccctat      900
agggggaagat tcattactta ggcgcatagc agaactatct gttactttta gctgcatcaa      960
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<210> 328
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-142-315 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-142-315.mis1, potential

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282

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<220>
<221> misc_binding
<222> 502..521
<223> 12-142-315.mis2, potential complement

<220>
<221> primer_bind
<222> 187..205
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 637..657
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-142-315 potential probe

<400> 328
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tatgagggca gagagaggag gggctcgccc ccagggccac acaggggtcta gagtaggcca      120
gggatgggag ttacttctgg ggtcctagtc actccggagc cgcctcact ctagtcccct      180
ttgtaaaaag gagatagaca ttgactgatt cactgacaac ctcaaccaag tccccaggaa      240
gaggagggag aaaccagag atgttacgac atctttccgg gggtatcttg ccaccagggt      300
ccaggcacccg atgtggccct gccagctct tggcaaacgt caggcagggg tggggccgcc      360
tccacgacct ggcactgtct ctctctgca ggctgtgcat gcagcacgtg gagggccaca      420
ggcagaggct ggccgagctg gtgctcaggg gcatggactc agaagtcctg agctgccgca      480
tcagcagcac agcggctctgc rtggaagtga ggcaccgggt gtgggcgggg tgcaagcgga      540
ggggatgggg tccctggatt cttctgggga tggcttgtct ccaaggcatg ggagtcgggg      600
ggacaggcga gcgccctcc tctgagtgtc cattctctca tctgtcctt cattcattca      660
tcacatgagt agtgaggggc aggcctgtct ggaaagacag gctgagccct gcagcattgc      720
agttaactga tgtgtgaccc aggcaagtac agtgacccag gggaccaggg gagggcacct      780
gggttctcgc tggggcaggc ttcctgcagg aggactcaga gaggccgagg ggctttgcta      840
ggtgggaagg cagggccccag cgaccctatc tcagagttag tggtggagcg tcttgagggc      900
aatgatgcc aatgcaccct aggaaccaac cacagcccag gccagaggcc ggccagtgtt      960
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<210> 329
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-142-321 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-142-321.mis1, potential

<220>
<221> misc_binding
<222> 502..520
<223> 12-142-321.mis2, complement

<220>
<221> primer_bind
<222> 181..199
<223> upstream amplification primer

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<220>  
 <221> primer\_bind  
 <222> 631..651  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-142-321 potential probe

<400> 329  
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 ggcagagaga ggaggggctc gccccaggg ccacacaggg tctagagtag gccagggatg 120  
 ggagttactt ctggggctct agtcactccg gagccgccct cactctagtc ccctttgtaa 180  
 aaaggagata gacattgact gattcactga 'caacctcaac caagtcccca ggaagaggag 240  
 ggagaaaccc agagatgtta cgacatcttt ccgggggttat cttgccacca ggggccaggc 300  
 accgatgtgg ccctgcccag ctcttggcaa acgtcaggca ggggtggggc cgctccacg 360  
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 ggctggccga gctgggtgctc aggggcatgg actcagaagt cctgagctgc cgcacagca 480  
 gcacagcggt ctgcatggaa rtgaggcacc ggggtgtgggc ggggtgcaag cggaggggat 540  
 ggggtccctg gattcttctg gggatggctt gtctccaagg catgggagtc ggggggacag 600  
 gcgagcgccc ctccctctgag tgctcattct ctcatctgct ccttcattca ttcacacat 660  
 gagtagtgca gggcaggccc tgctggaaag acaggctgag ccctgcagca ttgcagttaa 720  
 ctgatgtgtg acccaggcaa gtacagtgc ccaggggacc cagggagggc acctgggttc 780  
 tcgctggggc aggcttcctg caggaggact cagagaggcc gaggggcttt gctaggtggg 840  
 aaggcagggc ccagcgaccc tatctcagag tgagtgttg agcgtcctga gggcaatgat 900  
 gcccaatgca ccctaggaac caaccacagc ccaggccaga ggccggccag tggtttctgt 960  
 aaaggactgc agagtaaatt tttggctttg tggccaactc a 1001

<210> 330  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-143-453 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-143-453.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-143-453.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 50..68  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 533..552  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513

284

&lt;223&gt; 12-143-453 potential probe

&lt;400&gt; 330

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gcccaggatg agtatcaggg ccattctacct ggctatccgg gtagtcaaga acaccatctc 60
tgataccccg tccaaggtaa cagggcagag acctggaaat ggggggcact caggggaccc 120
tgccagggag gcaaaggga gtctgggctc tgggaggcag aggcttttga tggaaaaccc 180
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caagggtttt gtgaggtctt tatcgcatgt gaaaaaagca actaaggggt actcctctac 420
ccatcctcaa aaggctgaaa accacagtc caggtccat ggcagccaca tccaccttgg 480
agtggagctt ggttctggcc rtgggctggg gaaggaggt ccggcatagt gaggggtctg 540
cacagcagtt ttgaggaaca tcagagaggg aatggaggag aatgggggct agagatttag 600
gcagagctgt cttcagtcctc tggaggctgt ggagggaagc gagaagaggc tcgttctgca 660
tctccttcac atagtgatc tcatcaatgg agcaagcccc agggggtcca ggataacaga 720
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ccttctagcc cagagcacc ccaattccta gctctctatg tcacaggatt ggggtgccttg 960
gagccacccc caaccccag catgtccacg gtcctgtccc c 1001

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&lt;210&gt; 331

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-144-169 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-144-169.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-144-169.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 651..669

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 148..167

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-144-169 potential probe

&lt;400&gt; 331

```

cctgacctcg tgatccgcc accttgccct cccaaagtgc tgggattaca ggcgtgagac 60
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cctcacaacc tcacagtgtt tgagttcatg ccatacatgg gcatcaccct ggctaccata 300
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285

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tgtcccactt gagcctgcag gtctggctcg atcagggcgc cggagggaga ctgctgggta 420
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&lt;210&gt; 332

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-144-33 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-144-33.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-144-33.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 515..533

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 12..31

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-144-33 potential probe

&lt;400&gt; 332

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gctaagccag ctctggactg ttaggggttg gaggtggagg gcagcagggt tgcctctct 60
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286

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<210> 333  
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<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-146-174 : polymorphic base T or C

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-146-174.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-146-174.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 655..674  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 271..291  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-146-174 potential probe

<400> 333  
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ggatgcagtg attatttcca ctagaactgc tatatcatga ccatgaattt tgggggaatt 180  
tttttgagat ctgagttctc ttcacctcct tattctcttt ttgacactgg attcttttct 240  
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aaccaaaagaa cattctaaga tttcctatag ggtattaggt ctaatgggga tgtgttatgt 360  
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ggcacttggt aagcacacaa tgaacagtca tagaaagctg gccgagggta gagttcagtg 480  
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ttagacaggg gggctggggg ctatcccaga gttttgagag caaggcagag gactctgaat 660  
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gacaggggcc ctgaaatggg accatgacag ctgggtctga gagacagtgg tagaaacatc 840  
cagattcagc acttacttgc tggcttgat gcagggtcta gaacgaaaag agaagaggag 900  
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<210> 334  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>

287

<221> allele  
 <222> 501  
 <223> 12-146-47 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-146-47.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-146-47.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 528..547  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 144..164  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-146-47 potential probe

<400> 334  
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 attgtgaggg aatacactag taaaggtcac tcagttctaa ggggaaaaatg attaaccaaa 180  
 gaacattcta agatttccta taggggtatta ggtctaattgg ggatgtgtta tgtcaccaga 240  
 acaaacttct aagtttatat agcctctagt gacataacct gagaccgga cttggcactt 300  
 ggtaagcaca caatgaacag tcatagaaag ctggccgagg gtagagttca gtgtgaacaa 360  
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 ggggggctgg ggtctatccc rgagttttga gagcaaggca gaggactctg aattttcttc 540  
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 gagcagctga ggcagtgatt cagaagggac agctgggggt tgggcaactg ggggacaggg 660  
 gccctgaaat gggaccatga cagctgggtc tgagagacag tggtagaaac atccagattc 720  
 agcacttact tgctggcttg gatgcagggt ctagaacgaa aagagaagag gagtcacttc 780  
 tatacagaaa catgtccaga gtgcttactg tctgcaaaac tgtggactgg cacctgagtg 840  
 atagcatgat tccaaagcca aaatcttgcc tgtaaggaat atatgtatag gatatagcta 900  
 tactctaata gcaaggacag atatgcaaac tgctaaaaga tacaaggcag aacagaacaa 960  
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<210> 335  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-148-283 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-148-283.mis1, potential

288

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-148-283.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 765..783  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 375..395  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-148-283 potential probe

<400> 335  
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 ggattgactt ggagaagagc ctgaccataa tcttcaggat gaaataaagg cctggatgac 180  
 tgaaataaag actggagcct tcggcattca gaagagggaat tcagactgtg caagatctga 240  
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 agcctccctg cagctttttc tatattgaca gccacttcag agagagtcct ctctgaccc 480  
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 cagagaaaagc tggcttagca atgttgtagt gtttttggat gcgctgctt actcatacat 600  
 gagaagaaaag tttcaagggt tagcaaatag ggtcacatcc agcagagagc atatgactgg 660  
 gggctagggc aatggtgact cctcagacct cagctgctgc ctgataaaca tgggttaacag 720  
 agaagttaga gacagtgaca tgaaatgggt gttcacagcc ttgtgttggg aattggatga 780  
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 aatggttaat aattaactag aggagggcac tctgtcttcc aattacacgt tgatttgcta 960  
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<210> 336  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-148-311 : polymorphic base T or C

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-148-311.mis1, complement

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-148-311.mis2, potential

<220>  
 <221> primer\_bind  
 <222> 793..811  
 <223> upstream amplification primer, complement

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<220>
<221> primer_bind
<222> 403..423
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-148-311 potential probe

<400> 336
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ggaaccattc ttatcagaac ttggtgctgg attgacttgg agaagagcct gaccataatc 180
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aagaatctga tcatcacatc ttgagctcag cctccctgca gctttttcta tattgacagc 480
cacttcagag agagtcctct ytgatcctta caagaaatat cctggtgcga aaaacgacca 540
aaaccacata gccagcctcc acgctgttca gagaaagctg gcttagcaat gttgtatggt 600
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tcacagcctt gtgttgggaa ttggatgaga aacaagagct tgaacttgga tgttccccag 840
agtgcagaca gggctcagacg tgtttttcaa gatagtcacg atcggtcttt tccagggtgg 900
ggccacagt gaaaaacagt gatagattaa tgggttaataa ttaactagag gagggcactc 960
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<210> 337
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-149-320 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-149-320.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-149-320.mis2, potential complement

<220>
<221> primer_bind
<222> 800..820
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 368..387
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513

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290

&lt;223&gt; 12-149-320 potential probe

&lt;400&gt; 337

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cagcacctgt aattccagct aattgggagg ctgaggcagg agaattgctt gaaccggggt      60
caggggggttc ggaggtcgga ggttgccagt agtccggatc atgccactgc attccagcct      120
gggtgacaca gccagactct gtctcaaaaa caacaacaac aacaaaacaa caacaacaac      180
aacaacaaaa atctcactgg acatcctagt agctaaggct tccacatat tcatgattac      240
ttctgttggg aagtgcctta caacaaattg ctagttgtct cagtctgggt tcccctgaga      300
tgaggattca agggccagga gtttatntag gaagtaaagg aaacactgat agaggagtgg      360
cagagtgaga aggggtgatg gtcattccaca gctggctctc ttgtggtcaa tcggagctta      420
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ggaacacggc tgttgggtgc ytgagtactt gcctcgtcag ggattgaaac gtactcccag      540
gtagtagtaa tttctctgcc cttccattag gccacaaagg gggctctgac agagagagct      600
gacgagaaaa aacacacgcc cttgtcactg aagaggta caaggggatc gtgtggggca      660
ccacctgcac tgctaccctg gacaaatagc ttaagaaatc cccacactgc atcccaaac      720
ttactatcag cgtgtgaggg agacaggctc ccacaccctc attagcaca agtactatct      780
tgaaaaagaa agcctgtcag tttgatagga gaaaagcagg atcttgttta caatgtgctt      840
ttattattgt tattattaga gattgtattt cttttcaagc tgatgagccg tctgtgttta      900
ttttttggag gatacccttt gcccactttc ctattggagt gtattaccct gaggatttgg      960
taagagtgtc tattgcattc accagaatgt cttttttgtc a                                1001

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&lt;210&gt; 338

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-151-174 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-151-174.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-151-174.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 328..345

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 827..845

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-151-174 potential probe

&lt;400&gt; 338

```

gcctgtgtct ctgacatgca cagaaaaatt cctgtgtcta tctggaagat gagggaacac      60
ttatgaaatc aagtctctgc ccagagggc cccttgggag aggctgcagt gacctagtgc      120
cctgggctgg gtcaggagg gagagggtgg gattgtggtt ggcagagctc caggctttgt      180
ggggcctgtg gcttctgtag cggtgagagc cttttttaag aaaaaagata catagttagt      240
aaccacacat tggctaggaa aatgaatatt tactttgaga aaatcaagtg ctgcaaaatc      300
ataaattttg caagcttga caagtaccgc aacccttaaa atgtctagaa cgacgtagt      360

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291

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tttttactga ttgaccatct gacacagcct gatgagactt aattgtctac actttttggc 420
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ggtctggggc gcagggatgg kcgctctggtg caggagagaga agtgggctgg gcaggaggac 540
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caagtccaac actcagtaac atgtgaggcc agaggctgtg tgggaaggctt cacctgggct 660
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&lt;210&gt; 339

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-151-196 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-151-196.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-151-196.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 306..323

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 805..823

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-151-196 potential probe

&lt;400&gt; 339

```

gaaaaattcc tgtgtctatc tggaagatga gggaacactt atgaaatcaa gttctgtccc 60
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gaggggtggga ttgtggttgg cagagctcca ggctttgtgg ggcctgtggc ttctgtagcg 180
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tgaatattta ctttgagaaa atcaagtgtc gcaaaatcat aaattttgca aagcttgaca 300
agtaccgcaa cccttaaaat gtctagaacg acgtagtgtt tttactgatt gaccatctga 360
cacagcctga tgagacttaa ttgtctacac tttttggctc catgctgttt gactgtcttg 420
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gaggtgggga tgaaggctca gcaaaatgag gcaggggcca agtccaacac tcagtaacat 600
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caggaacaac tgggccc aag ttcagagctg tgaactgagc agagctgagt cagagggagt 840
cctgatccag ggcccacaga tccaggctgc tcagcctcca ggggcccagc tgtgggcagg 900

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292

ggaggctgat gcttggacag gggacagaca gtggcaccct ccccttcccc aaatgggtgt 960  
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<210> 340  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-151-270 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-151-270.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-151-270.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 232..249  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 731..749  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-151-270 potential probe

<400> 340  
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 taagaaaaaa gatacatagt tatgaacccc acattggcta ggaaaatgaa tatttacttt 180  
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 cccaagttca gagctgtgaa ctgagcagag ctgagtcaga gggagtcctg atccagggcc 780  
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<210> 341  
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 <212> DNA  
 <213> Homo Sapiens

<220>



293

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<221> allele
<222> 501
<223> 12-152-453 : polymorphic base C or T

<220>
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<222> 481..500
<223> 12-152-453.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-152-453.mis2, potential complement

<220>
<221> primer_bind
<222> 50..68
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 553..572
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-152-453 potential probe

<220>
<221> misc_feature
<222> 260
<223> n=a, g, c or t

<400> 341
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cctgcgaggg aggggcgtgg agcctggcac ccgtgggact catgggaatg aacactgccc      180
ttccgggttg aagggaccag gtgctgggtcc ctccggaggct gcagtgaggc tgtggccttt      240
gcattagttt gttagggttn gctgtaacaa agaaccacac atgcgtagct tgaagcaaca      300
gagatgtatt ctctgtacgt cctggaggtc aggagtccaa actcagggtg tcggcagggc      360
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tggtatggggc tgggtgccat gtaaagggat cactcatcta taaggagggt gtgggtctca      780
gctcttaaag caaggagtgg aaagggatgt gactgattta ggagccatct gtagatcatt      840
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cctagcgtag cacttcccaa gccttgctgt gggttagaac aacctggggg ctgggtgaaa      960
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<210> 342
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-153-116 : polymorphic base C or T

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294

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-153-116.mis1

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-153-116.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 386..405  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 867..887  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-153-116 potential probe

<400> 342  
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 cccttgacct cgtctctgtc cttgaaggag agggaccag acacccccga gtgctcagtt 180  
 ctcacacaca caggcacgca gagtaaaaat gaaggtgctg tgggggggtg gaagggcagg 240  
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 acccctgtgat gactgaggag gagtttggcc tgaaggtgtt ccccatgtat cgctacttcg 480  
 tgacagtgtg gctgaggcac yacaaccccg aggtgagatg caccctctt aggagggcct 540  
 ggtgggtccc aggccacagc tccccagcct gtggggcccg gaggtgctga gggaccgtcg 600  
 ggtccacact tactcccagg accctagagg gcagtgcct gactcatggc agagccgggg 660  
 aactcatgga tgctgaacag ctgggaccta ggagcctcat ccacgaggcc aggcttgatg 720  
 aaggttggtg ataactctga gagcacttgt tacatgctgg gcactctacc agcgtcttac 780  
 ccatgggagc tattttcaca cccactctgt gatataggac tagttgacaa atgaggaaac 840  
 tcaggcacag aggggttaag tacctgctca aggtcatgtg gctgctaagt gggagggact 900  
 caaaccggc cgctgtgca ggaaagacct tgctctgctt ctgaggcgcc ttctgtcctg 960  
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<210> 343  
 <211> 1000  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-154-480 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-154-480.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-154-480.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 23..43  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 522..540  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-154-480 potential probe

<400> 343  
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 aaacccttaa tataatcaca tctataaagt ctcttccacc tgtaatggaa tatattccca 180  
 ggtttgcgca actacaatgt agatactttg tggggctgtg attctgccta ccacagacac 240  
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 tttaaattta atttttgttt ctgtcctccg tagtcttcta ttctccaggc ttcagagctc 360  
 tcagcttacc attcaattat ctcttttctc tccttatatt ccttttttca ttttttaaaa 420  
 acaatacttt caaagagcaa aaattttaga atttgatgag gtttattcta gtgaagtgtt 480  
 tatcgtttgt actttttgta ycctaaggaa gctttgttta cccaagata tagtttctac 540  
 attctcttaa aaacactaaa gagttccagt ttatatgttt agctctgtga gattgggaga 600  
 ggggagctag atcactcagg tcaggcttcc gggatgcctt tttctgtctc tggacgttgc 660  
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 tatctgtgca gccagagccc ctctcatct ccagaccctg gaagctgatg ccttgggcag 780  
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 ctgagggccc ctatgtctct gaaggcacca ccatttcca agatacatgg gcctccgcag 900  
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 ccactacatg gaaatagaac accactacat ggaaatagaa 1000

<210> 344  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-155-403 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-155-403.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-155-403.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 99..116  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 628..647

296

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-155-403 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 49,58,62,639

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 344

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tnctaataag	gacacaacaa	agagtgaag	cattgctatg	tctattctgc	ttgcccagaa	120
tcttggctct	aaaaaatgaa	gagtgtttgg	gtgtggggag	gagcttcagt	gtgcatgtgc	180
atgcaaagta	cctactctaa	ggagaagaat	gagaggggtac	cctaattacc	tgtaatatg	240
tcccatagga	cacccaaaact	ctagtttagct	gtttctctat	gatcctctaa	gcacatcccc	300
aagtatggct	ggccagtgat	gtgtatgggt	caaatgttgg	gatctgtgca	gttatcttgg	360
aattgtatag	tacagcagta	tatccccccc	aaaaagagt	taatacttcc	aattctggct	420
gcacaatact	tgccccatag	tccatggtca	ataaatataaa	atttgagttg	tttttgctca	480
tctttccctt	ttgacttcaa	mtcagtcac	agaatttccc	caaatgcctt	tcccctggat	540
cttggggccag	tggaatgagt	acaatttaac	ttaattgaat	ttgcttatct	atttggttct	600
ctgttgtgaa	caaaagttct	ctgaaaagga	atttggaana	aagagacttt	gttctagtga	660
acagtttgca	aaccagggag	ttacagcctc	tggtacgcaa	tgaagggtgag	ttccacagaa	720
cacaaggcag	gcagggttca	cggcaaaaag	ttccttccca	ggttcccaat	caggtccatt	780
tatgcaaatg	aaggatggaa	acttgcttag	ttcttattgg	tcactgcagc	tgcatcttga	840
ttggttgatg	aagctgagcc	ctgagtggtc	gaggtgggtg	agctttaatt	ggttggttca	900
ggtgagcgct	gaaaatctca	actataaaaa	ggtacaggtt	ttcaggatac	tcagagtaac	960
cgtgtgacct	gtagtaagca	aagggccagt	tggtctctatt	t		1001

&lt;210&gt; 345

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-156-91 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 12-156-91.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-156-91.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 412..429

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 844..862

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

297

&lt;222&gt; 489..513

&lt;223&gt; 12-156-91 potential probe

&lt;400&gt; 345

```

cagctgtccg tgtcttctgc tgagatggcc acaggactcc aggttcccct gccgtggctg      60
gccacaggac tgctgcttct cctcagtgtc cagccctggg ctgagagtgg aaagggtgtg      120
gtggtgccca ttgatggcag ccactggctc agcatgcggg aggtcttgcg ggagctccat      180
gccagaggcc accaggcagt ggtcctcacc ccagaggga atatgcacat caaagaagag      240
aactttttca ccctgacaac ctatgccatt tcgtggaccc aggatgaatt tgatcgccat      300
gtgctggggc aactcaact gtactttgaa acagaacatt ttctgaagaa atttttcaga      360
agtatggcaa tgttgaacaa tatgtctttg gtctatcata ggtcttgtgt ggagctacta      420
cataatgagg ccctgatcag gcacctgaat gctacttcct ttgatgtggt tttaacagac      480
cccgtaacc tctgcggcgc rgtgctggct aagtacctgt cgattcctac tgtgtttttt      540
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tatattccta gattactaac aaccaattca gaccacatga cattcatgca aagggtcaag      660
aacatgctct accctctggc cctgtcctac atttgccatg ctttttctgc tccttatgca      720
agccttgccct ctgagctttt tcagagagag gtgtcagtgg tggatattct cagtcatgca      780
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ggtgccttca tccaatcaat gttccaggca aaacactttt taaaaaatg tatttattta      960
aaattgcttc catatttact tatctttcca aatatttcat t                                1001

```

&lt;210&gt; 346

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-157-437 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-157-437.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-157-437.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 67..87

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 513..533

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-157-437 potential probe

&lt;400&gt; 346

```

tcttttctat aattctcagc cagatttcaa gttgggtgct tctccttggga tttatggcac      60
atgggtgggc tgctcttcca ttgattgggg atattttgag caaaataaca gtgataacac      120
tgaagagcat tgcaagcatc cttcaagatg agtcaggctc agtgagtggg ttgaagggtc      180
cagtcaacta tttgttgata ccactgggtac aaaatgatct aaagcccaag acttaccttc      240
ctggagatgc caatccattg gtggggaggg ggaaggggga agaagttatg gatttataaa      300

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298

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attagatattt aaagagtttg ggtcatctct aagaatacct cccctatagg ggaagattca 360
ttacttaggc gcatagcaga actatctgtt actttaagct gcatcaaaca ggcagatttt 420
aaaataaaatg tcatgaacat caactaagca caatgatatg ctgagtactg aagggaacaca 480
gagatacata agacctagtc mcagcctttt aaggacttag agataagaca gacacataca 540
aacaggatta taggtcaaag ctgactaaca ggactgtgtt tacaggtaaa cccacacaca 600
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ttatacaaaa ggagctatca agaggcttgg tgcaaagtgt gcatttggct tttgacctgg 720
gcttaggcag aacgactgga cttggcgctt cctaggagag tgaatacaa gacactttgg 780
gggagcagga gtttaaagca ggatggtggc tctcaactct aggggcacaa tagaactcat 840
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tgttgatgaa ggagctagaa gaagaatgag ctagcatgca c 1001

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&lt;210&gt; 347

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-158-213 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-158-213.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-158-213.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 691..710

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 230..247

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-158-213 potential probe

&lt;400&gt; 347

```

ttttctgtct ctggacgttg ctggggtgac ctcaactgaca cccatggctt cagctaccac 60
atatgctgat ggctccaagt ctatctgtgc agcccagacc cctcctcatc tccagacctt 120
ggaagctgat gccttgggca gctcctccta tttcccaggc accagggaatg tgagcttctc 180
cctccccaca gtcctgctgt cctcagggcc cctatgtctc tgaaggcacc accatcttcc 240
aagatacatg ggcctccgca gggcttagga gtgccagaca cgtaaccaga aatcagatga 300
catcacatc taaataaaac accactacat ggaaatagaa caccactaca tggaaataga 360
acatgggagc cccttgaatg tggcaagagc accctcccag gcatgttcca ccctacccc 420
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agaagctgag atgttgacac yacagctgag aatccctttc tagcactctg tgcctcaca 540
aatccccaga aatcgtcctc ccctggggag ttctcaagcc cttacagacc tgccctctct 600
gtgccatcct gcatatgcct cccttgagct ggggtgtccct ctgatggacg catccattca 660
ctgcctgtcc catgggttgt gtccaaaggt ggaatctgtt atcaatgttg atttctaatt 720
ggagtaactt cctccataag ggaagcctca gcctcaccag caatggcaga catggccagg 780
catgtagaca cagaggtagt gagatggaaa gtgggcacag cccagagagc ctgcccacct 840

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299

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catcctgggg cgaccaggac aaggaagcat cagcaatctt gcgagcacat gtaggagtga    900
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cctagaaagg cacacattta attctccatt ttgaaatttg a                          1001

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&lt;210&gt; 348

&lt;211&gt; 980

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 480

&lt;223&gt; 12-158-450 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 460..479

&lt;223&gt; 12-158-450.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-158-450.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 907..926

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 446..463

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 468..492

&lt;223&gt; 12-158-450 potential probe

&lt;400&gt; 348

```

atactttcaa agagcaaaaa ttttagaatt tgatgagggt tattctagtg aagtttttat    60
cgtttgtact ttttgtaccc taaggaagct ttgtttaccc caagatatag tttctacatt    120
ctcttaaaaa cactaaagag ttccagttta tatgttttagc tctgtgagat tgggagaggg    180
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ggtgacctca ctgacacca tggcttcagc taccacatat gctgatggct ccaagtctat    300
ctgtgcagcc cagaccctc ctcatctcca gacctggaa gctgatgcct tgggcagctc    360
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ctaggagtgc cagacacgta accagaaatc agatgacatc actatctaaa taaaacacca    540
ctacatggaa atagaacacc actacatgga aatagaacat gggagcccct tgaatgtggc    600
aagagcacc ccccaggcat gttccaccct caccctgggc tcatcaggag ggttcttaag    660
atgcagacag ttttaagggg gttggaggaa tagttgagaa gctgagatgt tgcaccacac    720
gctgagaatc ctttcttagc actctgtgtc ctacaaaatc cccagaaatc gtcctcccct    780
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aaaggtggaa tctgttatca atgtggattt ctaatgggag taacttcctc cataagggaa    960
gcctcagcct caccagcaat

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&lt;210&gt; 349

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

300

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<220>
<221> allele
<222> 501
<223> 12-161-157 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 12-161-157.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-161-157.mis2, potential complement

<220>
<221> primer_bind
<222> 345..363
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 729..749
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-161-157 potential probe

<400> 349
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<210> 350
<211> 998
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 498
<223> 12-162-21 : polymorphic base A or G

<220>
<221> misc_binding
<222> 478..497
<223> 12-162-21.mis1, potential

```



<220>  
<221> misc\_binding  
<222> 499..518  
<223> 12-162-21.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 478..497  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 909..927  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 486..510  
<223> 12-162-21 potential probe

<400> 350  
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atacatgggc ctccccaggg tctaggagtc ctagacgtgg gagtagaaat tagatgacat 180  
tggtgtctaa acaaaacacc actgcatgga gatagaacag gggagccccc tgaatgtggt 240  
gagagcacc tcccaggtgt gttccaccct caccgcgggc tcatcaggag ggtgcttagg 300  
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aactagagcc cccacaaatt attctgttct ggaggaacca ttcttatcag aacttggtgc 900  
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<210> 351  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 503  
<223> 10-470-25 : polymorphic base A or T

<220>  
<221> misc\_binding  
<222> 484..502  
<223> 10-470-25.mis1

<220>  
<221> misc\_binding  
<222> 504..523  
<223> 10-470-25.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 479..498

302

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 880..899

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-470-25 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 271,364,831,842

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 351

```

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agtaactttt cctcagtgct cagagtcagg gaagtcaacc actaatgact tcaaactaaa      180
ataattctgt agaaaacctg cctaaaataa gcatatgtga tttagcgagc aacaatatag      240
cattaaagcc aactgggtgcc actttaaaga ncctatatta gtacttataa tatgataagt      300
gaagagtttg ggtatctcct caaatactgt gtaataactc tatttcattt ctccctttca      360
caancgcaca cacatacaca cacacatatt tacacaaaga cccttaacag aggcaaccta      420
tctcatatta tacatatgtc aaaaaaaaac tgagtaattg agtcagttaa aaaacatcct      480
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gtgcttgctg tgctgataat ctattatgat agaacaaatt cttttttttc acaggaaatg      600
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caaaaaggta agataaaatg ttttaatggt gtaaaaaact actgaaagag gctgttaaag      780
tttgtaaaga acccaaattg tagaaacttc ctgcctatat atttcagctg ntgggaaaag      840
cnctaattat ctcagatatt aattcaaaat caaaaatatg tatggaagat gataaactca      900
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acaattaaga gttgcaggta aagttttggt attatcatga t                               1001

```

&lt;210&gt; 352

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-471-84 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 484..502

&lt;223&gt; 10-471-84.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-471-84.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 420..439

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

303

<222> 788..807  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 10-471-84 potential probe

<400> 352  
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 tcagcctctc aagtagctgg gactacaggc acatgccacc acgcctgggt aatctttttt 240  
 tttttttttt ttttttttca tttttttact ggagacgggg tgacgggggt tcaccgtgtt 300  
 agccaggatg gtcttgatct cctgacctcg tgatccgccc gcctcgacct cgaaaagtgc 360  
 tgggattgca ggtgtgagcc tccgtgcttg gccaaattaa cttactttca atgttgatac 420  
 ttttctgctt atcgtttaga tataaagaga atgctatgaa attatcaaga attcatcatg 480  
 atcaaccagt gaagcccctt gawagagcag tcttctggat tgaatttgtc atgcgccata 540  
 aaggagccaa gcaccttcgg gttgcagccc acgacctcac ctggttccag taccactctt 600  
 tggatgtgac tgggttcctg ctggcctgtg tggcaactgt gatattcatc atcacaaaat 660  
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 gtctgaggct ggaagctggg aaaccaata aatgaactcc tttagtttat tacaacaaga 780  
 agacgttttg atacaagaga ttcctttctt cttgtgacaa aacatctttc aaaacttacc 840  
 ttgtcaagtc aaaatttgtt ttagtacctg tttaaccatt agaaatattt catgtcaagg 900  
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<210> 353  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 10-471-85 : polymorphic base A or C

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 10-471-85.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 505..523  
 <223> 10-471-85.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 420..439  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 788..807  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 10-471-85 potential probe

<400> 353

304

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agtgggtgga ttgtggctca ctgcaacttc cgcctcttgg gttcaagcga ttctcctgcc   180
tcagcctctc aagtagctgg gactacaggg acatgccacc acgcctgggt aatctttttt   240
tttttttttt ttttttttca tatttttact ggagacgggg tgacgggggt tcaccgtggt   300
agccaggatg gtcttgatct cctgacctcg tgatccgccc gcctcgacct cggaaagtgc   360
tgggattgca ggtgtgagcc tccgtgcttg gccaaattaa cttactttca atgttgatac   420
ttttctgctt atcgtttaga tataaagaga atgctatgaa attatcaaga attcatcatg   480
atcaaccagt gaagcccttt gamagagcag tcttctggat tgaatttgtc atgcgccata   540
aaggagccaa gcaccttcgg gttgcagccc acgacctcac ctggttccag taccactctt   600
tggatgtgac tgggttcctg ctggcctgtg tggcaactgt gatattcatc atcacaaaat   660
gtctgttttg tgtctggaag tttgttagaa caggaaagaa ggggaaaaga gattaattac   720
gtctgaggct ggaagctggg aaaccaata aatgaactcc tttagtttat tacaacaaga   780
agacgttggt atacaagaga ttcctttctt cttgtgacaa aacatctttc aaaacttacc   840
ttgtcaagtc aaaatttggt ttagtacctg ttttaaccatt agaaatattt catgtcaagg   900
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&lt;210&gt; 354

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-472-202 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 484..502

&lt;223&gt; 10-472-202.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 525..543

&lt;223&gt; 10-472-202.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 304..322

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 714..732

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-472-202 potential probe

&lt;400&gt; 354

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gatataaaga gaatgctatg aaattatcaa gaattcatca tgatcaacca gtgaagcccc    60
ttgaaagagc agtcttctgg attgaatttg tcatgcgcca taaaggagcc aagcaccttc   120
gggttgagc ccacgacctc acctgggtcc agtaccactc tttggatgtg actgggttcc   180
tgctggcctg tgtggcaact gtgatattca tcatcacaaa atgtctgttt tgtgtctgga   240
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ggaaacccaa taaatgaact ctttagttt attacaacaa gaagacgttg tgatacaaga   360
gattcctttc ttcttgtagc aaaacatctt tcaaaaactta cttgtcaag tcaaaatttg   420
ttttagtacc tgtttaacca ttagaaatat ttcatgtcaa ggaggaaaac attagggaaa   480
acaaaaatga tataaagcca taygagggtta tattgaaatg tattgagctt atattgaaat   540

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305

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ttattgttcc aattcacagg ttacatgaaa aaaaatttac taagcttaac tacatgtcac    600
acattgtaca tggaaacaag aacattaaga agtccactga cagtatcagt actgttttgc    660
aaatactcag catacttttg atccatttca tgcaggattg tggtgtttta actgttggtg    720
aggaagctaa taaataatta aattgtatag aaagtctctt cctcttgata ttttgagatg    780
attagtgtcg cttggccttt attgtgcac gtgcttcaac gtcatttttt ttcctaaaag    840
gtatgataaa aatgcttacc atttttagagc ttaagtcatt tcccagtgaa aagtatgtgg    900
aattagaaat atagcaactc ctacctgggt tctactacaa aatgaactaa ttttacaatg    960
cgtttggttt tttgagccaa ttctattttt ctgttcattt g                                1001

```

&lt;210&gt; 355

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-473-333 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-473-333.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-473-333.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 171..189

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 582..600

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-473-333 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 887

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 355

```

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tacaagagat tcttttcttc ttgtgacaaa acatctttca aaacttacct tgtcaagtca    120
aaatttgttt tagtacctgt ttaaccatta gaaatatttc atgtcaagga ggaaaacatt    180
agggaaaaca aaaatgatat aaagccatat gaggttatat tgaaatgtat tgagcttata    240
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atgtcacaca ttgtacatgg aaacaagaac attaagaagt cactgcacag taccagtact    360
gttttgcaaa tactcagcat actttggatc catttcatgc aggatttgtg tgttttaact    420
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tgagatgatt agtgcctgctt ggyttttatt gtgcacgtg cttcaacgtc attttttttc    540
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cttttttatt ctttgttttt taggtatttc aatagctttt gggtgacaac tgggttttgg    780

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306

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agtatacaact gtacccagtg tgtactgttt tatccctcac atccctncct acccttcctt      900
ctgagttcca agagtccatt atatcattct tatgcctcta tgctctaata gcttaggtcc      960
cccttataag tgagaacata cattgtttgg ttttatattt c                          1001

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&lt;210&gt; 356

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-494-284 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-494-284.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-494-284.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 220..238

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 624..641

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-494-284 potential probe

&lt;400&gt; 356

```

ctggtgtgag atggtatctc tttgtggatt tgaccagtga tgtaaaccct tttttcatat      60
agtggtttgc cacatatagt tttcttttga aaagtgtaac aactttttta atacttgaac      120
ttttcattga ttatcttatt tgtctaagct actattttga aaaatcatga tttccttata      180
tacctaatta tgaaattaag gaaatgaaat atgagtattc tatttacatc agtctgagta      240
gttcttggtta cttaacatcc cttgttcttc tcattgttaa tctctttaga tttctaacat      300
tctatgactt ttgagttcca ctcattggaat aagatatttt cttcactgta acagggttctg      360
tggagatttg atgggaataa accagatact ttaggactca atactcggct gtacaagtgg      420
ataccccgaga atgatcttct tggtaagtct ctgaagaaca aatactgaat atattagtaa      480
cagattatta aagtgttaat agytatcatg aaacaagctt actgaacatt tgttatggaa      540
aaacttaaaa tgaaacttct ttatatttat tttccagtcc cgggggaaaa gaataaatga      600
taattgttgg cattttatga tatgcaccca cattctttac aatcagagtc agagtatctt      660
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tccctaactt taatcaaaaa gtgatgacac atttcataat gaaatgtgac ctgtctttcc      900
tcaattctag caccaccacc acctcactgc ctgctgcctt gcacacccta catatcacac      960
tccgtgactg tacttaagag aacacattct ggctgggcac g                          1001

```

&lt;210&gt; 357

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

<220>  
<221> allele  
<222> 499  
<223> 12-637-219 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 500..519  
<223> 12-637-219.mis1, potential complement

<220>  
<221> misc\_binding  
<222> 480..498  
<223> 12-637-219.mis2

<220>  
<221> primer\_bind  
<222> 698..717  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 230..250  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 487..511  
<223> 12-637-219 potential probe

<220>  
<221> misc\_feature  
<222> 953  
<223> n=a, g, c or t

<400> 357  
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ttgtgataca agagattcct ttcttcttgt gacaaaacat ctttcaaaac ttaccttgct 180  
aagtcaaaaat ttgttttagt acctgtttta ccattagaaa tatttcatgt caaggaggaa 240  
aacattaggg aaaacaaaaa tgatataaag ccatatgagg ttatatgaa atgtattgag 300  
cttatattga aattttattgt tccaattcac aggttacatg aaaaaaaatt tactaagctt 360  
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agtactgttt tgcaataact cagcatactt tggatccatt tcatgcagga ttgtgttggt 480  
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tttttcctaa aagggtatgat aaaaatgctt accatttttag agcttaagtc atttcccagt 660  
gaaaagtatg tggaattaga aatatagcaa ctctacctg gtttctacta caaaatgaac 720  
taattttaca atgcgttttg ttttttgagc caattctatt tttctgttca ttgaaaata 780  
ttcatctttt ttattctttt gttttttagg tatttcaata gcttttgggt gacaactggg 840  
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ttccctctga gttccaagag tccattatat cattcttatg c 1001

<210> 358  
<211> 668  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 499

308

&lt;223&gt; 12-639-95 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-639-95.mis1, potential complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 480..498

&lt;223&gt; 12-639-95.mis2

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 573..593

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 144..164

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-639-95 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 57,198,240,552,616,630

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 358

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caaataaaaag	ttaaactactg	taaattattg	ttcagtttgt	gaggattgct	tcggagtgtc	180
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aaaattgtca	cttgaaattt	ttcttggaag	tagtgcttga	taatttgtaa	ttctaatagaa	480
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ggagatga						668

&lt;210&gt; 359

&lt;211&gt; 814

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-639-241 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-639-241.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding



309

<222> 500..519  
 <223> 12-639-241.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 719..739  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 290..310  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 487..511  
 <223> 12-639-241 potential probe

<220>  
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 <222> 38,95,116,203,344,386,698,762,776  
 <223> n=a, g, c or t

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 gcttgataat ttgtaattct aatgaagtta tacataaaac cctagcagca tttatttatt 660  
 tcttataagt agcttatttt atagacttgc tattttgnca atcaaaggac aacaggctct 720  
 aatataataa cctaccgaca aagtagatac atttgaactt cntctcttat tattcnaaaa 780  
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<210> 360  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 499  
 <223> 12-640-151 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 479..498  
 <223> 12-640-151.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 500..519  
 <223> 12-640-151.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 629..649  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 156..176  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 487..511  
 <223> 12-640-151 potential probe

<220>  
 <221> misc\_feature  
 <222> 62  
 <223> n=a, g, c or t

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 aatgagactc ccaagactga ttcataaaaa ttccaaatca caataactaga ctcaggaatg 180  
 tcagtgtatc ttaaccacca gcttttattt tcattttttg aaaaactact ggaaaactct 240  
 gacaaacttt aagtgaagca taaagcattg tagaggaaca taaatgtaga tataaaaatta 300  
 tcccaactgt gaatagcttt tcctcagtgc tcatatttag ggaagtagac cactaatggc 360  
 ttcaaaactaa agaatttcta cagaaaacct gcctgaaata aacacaagtg atttagtaga 420  
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 atatttacac aaatatcctt aacagaggcc aactatctca aatatcttct tgcaaagaaa 660  
 ctgagtgatt gagtcagtta aaaaatatta tttactccaa taattcctca aaataactga 720  
 ttttctctct ttaatatattg gtaccagttc tttagtagtg cctgctgtgg tgatactctt 780  
 ttgtgattaa acaatttttt tttcacagga aatggaggag tttgtacaga gctctggaga 840  
 aaatgggtgt gtgggtgttt ctctgcggtc aatcataagt aacatgacag cagaaagggc 900  
 caatgtaatt gcaacagccc tggccaagat cccacaaaag gtaagataaa gtaccttact 960  
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<210> 361  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 499  
 <223> 12-640-296 : polymorphic base T or G

<220>  
 <221> misc\_binding  
 <222> 479..498  
 <223> 12-640-296.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 500..519  
 <223> 12-640-296.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 769..789  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 296..316

311

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-640-296 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 1..4,6..8,12..22,25..28,30,32..37,39,41..44,47..50,202

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 361

```

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agagttccct acatgcagtt agaaatagca catcaattta acagtgtgat ttcagggcaa      180
taggtgttcc acctaaacaa tnaacctgaa aggtacaatt attcaacaac taactataaa      240
ctctacaatt ccatgtgata aatgagactc ccaagactga ttcataaaaa ttccaaatca      300
caataactaga ctcaggaatg tcagtgattc ttaaccacca gcttttattt tcatTTTTTg      360
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aacacaagtg atttagtaga acaaaaatat aggattaaag cctagtgggtg ccacttttcc      600
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aatatcttct tgcaaagaaa ctgagtgatt gagtcagtta aaaaatatta tttactccaa      840
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cctgctgtgg tgatactctt ttgtgattaa acaatttttt tttcacagga aatggaggag      960
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&lt;210&gt; 362

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-640-325 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..518

&lt;223&gt; 12-640-325.mis1, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-640-325.mis2, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 797..817

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 324..344

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

312

&lt;222&gt; 487..511

&lt;223&gt; 12-640-325 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 6,24..25,28..32,34..36,40..50,53..56,58,60..65,67,69..72,75..78

230

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 362

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nnnnnnanann	nncnnnnnga	agaaaggcct	ggtggcctct	tctattcttg	tgccagtgtc	120
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tcaatttaac	agtgtgattt	cagggcaata	ggtgttccac	ctaaacaatn	aacctgaaag	240
gtacaattat	tcaacaacta	actataaact	ctacaattcc	atgtgataaa	tgagactccc	300
aagactgatt	cataaaaatt	ccaaatcaca	atactagact	caggaatgtc	agtgattctt	360
aaccaccagc	ttttattttc	attttttgaa	aaactactgg	aaaactctga	caaactttta	420
gtgaagcata	aagcattgta	gaggaacata	aatgtagata	taaaattatc	ccaactgtga	480
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gtcagttaaa	aaatattatt	tactccaata	attcctcaaa	atacttgatt	ttctctcttt	900
aatatttgg	accagttctt	tagtagtgcc	tgctgtggtg	atactctttt	gtgattaaac	960
aatttttttt	tcacaggaaa	tggaggagtt	tgtacagagc	t		1001

&lt;210&gt; 363

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-640-413 : polymorphic base C or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-640-413.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-640-413.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 885..905

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 412..432

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-640-413 potential probe

313

<220>  
<221> misc\_feature  
<222> 94,112..113,116..120,122..124,128..138,141..144,146,148..153,155  
157..160,163..166,318  
<223> n=a, g, c or t

<400> 363  
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acaattccat gtgataaatg agactcccaa gactgattca taaaaattcc aaatcacaaat 420  
actagactca ggaatgtcag tgattcttaa ccaccagctt ttattttcat tttttgaaaa 480  
actactggaa aactctgasa aactttaagt gaagcataaa gcattgtaga ggaacataaa 540  
tgtagatata aaattatccc aactgtgaat agcttttcct cagtgtctcat atttagggaa 600  
gtagaccact aatggcttca aactaaaaga attctacaga aaacctgcct gaaataaaca 660  
caagtgtatt agtagaacia aaatatagga ttaaagccta gtgggtgccac tttccaaga 720  
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<210> 364  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 499  
<223> 12-641-120 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 479..498  
<223> 12-641-120.mis1, potential

<220>  
<221> misc\_binding  
<222> 500..519  
<223> 12-641-120.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 600..618  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 127..147  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 487..511  
<223> 12-641-120 potential probe

<220>  
<221> misc\_feature  
<222> 5,519,564,828,846..847,850..854,856..858,862..872,875..878,880

314

882..887,889,891..894,897..900

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 364

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actttcagcc	ataatacaaa	caaagtgaat	tagaaaaatga	taaatatagc	aagatggcaa	180
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&lt;210&gt; 365

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-641-122 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-641-122.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-641-122.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 602..620

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 129..149

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-641-122 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

<222> 7,521,566,830,848..849,852..856,858..860,864..874,877..880,882  
884..889,891,893..896,899..902

&lt;223&gt; n=a, g, c or t

315

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acacttttcag ccataatata aacaaagtga attagaaaat gataaatata gcaagatggc    180
aagttttttgc aggaggaaat gtatatacaa taagattaaa attttctaaa taagtgtcac    240
atgtatacat tgacctatat aaatttgaca aaatccataa taaaaaatac atatttttcta    300
aaaattatac ctctagatag aaattttaga aaaattatct tttaaaaagg ttttcatact    360
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acactctgcc atatgtgat cactgtttac atttagactt ttttttcttt gttttgttaa    720
aaaccatttg gaaaagtttt accccaatga ttaaactctga aaatatattaa atttaaatat    780
tgggtatacat tgggggaactc aagtcagaat aattctcaat caattgcagn caagcacatc    840
ttcaccanna gnnnnntnnn gtgnnnnnnn nnnnctnnnn anannnnnna nannnnccnn    900
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<210> 366
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 432
<223> 12-641-223 : polymorphic base G or A

<220>
<221> misc_binding
<222> 412..431
<223> 12-641-223.mis1, potential

<220>
<221> misc_binding
<222> 433..452
<223> 12-641-223.mis2, potential complement

<220>
<221> primer_bind
<222> 636..654
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 163..183
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 420..444
<223> 12-641-223 potential probe

<220>
<221> misc_feature
<222> 24,41,555,600,864,882..883,886..890,892..894,898..908,911..914
916,918..923,925,927..930,933..936
<223> n=a, g, c or t

<400> 366
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ttagaagtaa tccaacatct ttatacgaaa gaatgattaa aacttatgtg catatagtag    120

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316

cattaaaata	atTTtatcaa	tgaaatgttc	agttacactt	tcagccataa	tacaaacaaa	180
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acaataagat	taaaattttc	taaataagtg	tcacatgtat	acattgacct	atataaattt	300
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tagaaaaatt	atctttttaa	aagggtttca	tacttttata	aaatttctta	cattaatttt	420
gtattatttt	trtaatatTa	tcacaaaagg	agaaacaagc	agtccatatc	agtgatgaat	480
ctctaaacag	aggtttagatt	ccttcaacac	ctggtagaaa	aacaaggctc	tggtgtcaag	540
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gttaacgtaa	gattaaaaat	catattttta	gaaaacactc	tgccatatgt	gatgcactgt	720
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atgattaaat	ctgaaaatat	ttaaatttaa	atattggtat	acattgggga	actcaagtca	840
gaataattct	caatcaattg	cagncaagca	catcttcacc	annagnnnnn	tnnngtggnn	900
nnnnnnnnct	nnnnanannn	nnnanannnn	ccnnnnngaag	aaaggcctgg	tggcctcttc	960
tattctggtg	ccagtgtctg	ctctgagaca	caacaaagtg	a		1001

&lt;210&gt; 367

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 388

&lt;223&gt; 12-641-267 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 368..387

&lt;223&gt; 12-641-267.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 389..408

&lt;223&gt; 12-641-267.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 636..654

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 163..183

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 376..400

&lt;223&gt; 12-641-267 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

<222> 24,41,555,600,864,882..883,886..890,892..894,898..908,911..914  
916,918..923,925,927..930,933..936

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 367

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cattaaaata	atTTtatcaa	tgaaatgttc	agttacactt	tcagccataa	tacaaacaaa	180
gtgaattaga	aaatgataaa	tatagcaaga	tggaagttt	ttgcaggagg	aaatgtatat	240
acaataagat	taaaattttc	taaataagtg	tcacatgtat	acattgacct	atataaattt	300



317

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gacaaaaatcc ataataaaaa atacatatatt tctaaaaaatt atacctctag atagaaattt 360
tagaaaaaatt atctttttaa aagggttyca tacttttata aaatttctta cattaatttt 420
gtattattttt tataatatta tcacaaaagg agaaacaagc agtccatatt agtgatgaat 480
ctctaaacag aggttagatt ccttcaacac ctggtagaaa aacaaggctc tgggtgcaag 540
ccagtgcgat gaaanactg aaaacgagtt acattaaatg tggctacagg taactgcaan 600
tcacatacaa cacctagaag ccccggtat tacgtggaat agtagtaca ggactctcac 660
gttaacgtaa gattaaaaat catattttta gaaaacactc tgccatatgt gatgcactgt 720
ttacatttag actttttttt ctttgttttg ttaaaaacca tttggaaaag ttttacccca 780
atgattaaat ctgaaaatat ttaaatttaa atattggtat acattgggga actcaagtca 840
gaataattct caatcaattg cagncaagca catcttcacc annagnnnnn tnnngtgnnn 900
nnnnnnnnct nnnnanannn nnnanannnn ccnnnngaag aaaggcctgg tggcctcttc 960
tattctggtg ccagtgcctg ctctgagaca caacaaagtg a 1001

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&lt;210&gt; 368

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 12-642-387 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 12-642-387.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 12-642-387.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 117..135

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 592..612

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 12-642-387 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 537,804,808,881,899,903,920,936,958,962..963,967,985,995

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 368

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tttattcgaa tattttactc tcattaaaca ttaagtaata tataattata attttagaaa 60
acctaaaata actcaatatt aatgcttatt tatgtatatt tctaaactga aaaaaaacg 120
agattcactg aaaactcagg ctggttattt gagaaggcca gaaatagtgt gtttctactg 180
tgcattttta gaaaaccctt ttttattttt attttatttt ttatttccaa catttattgt 240
aaattcagag gtacatgtgc aagaggtata ggtttggtac ataggtaaaa aaacactctt 300
gtataaacia atacaatgat atggactata tgtgtccctg tcaagaaaaa cagagcaaaa 360
cccttcttaa aaatggcaag acacatttta ttctgactgg tgaattaggg gatagagact 420
acagtataaa ttgagctcaa ctctaattaa gacaaaggta actggggctt ttaaagagaa 480
aaccgatatg actaaaaatg acraaaacat aaaggagcta ggaggttata tgaaaantgg 540

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318

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aatattacat aaaaattcag tgtgaaataa ttgaaaatta ttaggtaagg tgggttaggc      600
agtgtatatt ttgcagtttg gcaggatcat attcctttga gccaaagacct accatggaag      660
ctagggtggc ctttaaagac aaattcatga cctagtgtccc atataagcta agctaaactt      720
ggcccagtcct cttaacatgg tgtttatgca agtcttttgt gttacgtggg agtgtaaaat      780
tcaagtcctt catgaaatac aaanatantt tcttaatggg aatattctgt gtttcaaaat      840
tattaacggt cgtaactgac ttcctatcca aaactgtgca ncctggtaaa aatccttgnt      900
tanaaaagaca aaagatgtn tccacatgag gcctgnaata tgcttgagtt actatttnca      960
cnntagnaac aaccctttgt aatgngataa taatncacag a                          1001

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&lt;210&gt; 369

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 12-642-417 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 484..502

&lt;223&gt; 12-642-417.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 12-642-417.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 87..105

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 562..582

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 12-642-417 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

<222> 507,774,778,851,869,873,890,906,928,932..933,937,955,965,976,981  
983,988..989,998,1001

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 369

```

ttaagtaata tataattata attttagaaa acctaaaata actcaatatt aatgcttatt      60
tatgtatatt tctaaactga aaaaaaacg agattcactg aaaactcagg ctgtttattt      120
gagaaggcca gaaatagtgt gtttctactg tgcattttta gaaaaccctt ttttattttt      180
attttatttt ttatttccaa cattttattgt aaattcagag gtacatgtgc aagaggata      240
ggtttggttac ataggtaaaa aaacactcct gtataaacia atacaatgat atggactata      300
tgtgtccctg tcaagaaaaa cagagcaaaa cccttcttaa aaatggcaag acacatttta      360
ttctgactgg tgaattaggg gatagagact acagtataaa ttgagctcaa ctctaattaa      420
gacaaaaggta actggggcctt ttaaagagaa aaccgatatg actaaaaatg acgaaaacat      480
aaaggagcta ggagggttata tgraantgg aatattacat aaaaattcag tgtgaaataa      540
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attcctttga gccaaagacct accatggaag ctagggtggc ctttaaagac aaattcatga      660
cctagtgtccc atataagcta agctaaactt ggcccagtcct cttaacatgg tgtttatgca      720

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319

agtcttttgt	gttacgtggg	agtgtaaaat	tcaagtcctt	catgaaatac	aaanatantt	780
tcttaatggg	aatattctgt	gtttcaaaaat	tattaacggg	cgtaactgac	ttcctatcca	840
aaactgtgca	ncctggtaaa	aatccttgnt	tanaaagaca	aaagatgttn	tccacatgag	900
gcttgnaata	tgcttgagtt	actatttnca	cnntagnaac	aaccctttgt	aatgngataa	960
taatncacag	aataanagaa	nangatantt	ccctttantt	n		1001

&lt;210&gt; 370

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 434

&lt;223&gt; 12-646-429 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 414..433

&lt;223&gt; 12-646-429.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 435..454

&lt;223&gt; 12-646-429.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 6..26

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 474..494

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 422..446

&lt;223&gt; 12-646-429 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 31..32,35,59,514

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 370

ctacacttgg	aaaatatctg	taccccaa	at	nnganagtca	atataacctg	gtgggtaana	60
gctggcactc	tggatccaga	tggccttgat	tcaagtcctt	gaaaccactg	tataatgatg		120
tgggactgga	caagttactt	gaacttcatt	tgactcagct	tcctcaccaa	tgaaatggca		180
atagttgtca	tcacattatc	aatatcatgg	gttgctgaaa	gaacgaattc	attaatattg		240
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caaaaaattt	ccacaataat	aatttttagtg	ttcatataat	tgcataatat	attatagttt		360
caaaacatgc	tatgttttcta	atgcaaata	gac	agttatttact	atttcaagca		420
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ttgttgtag	ttctctccaa	ttttatattg	aagtattttc	ttgagaatgg	ccataaacca		960

320

agatcctgga aagctctgaa taaggtgggg aagtgagttt c

1001

&lt;210&gt; 371

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 438

&lt;223&gt; 12-646-433 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 418..437

&lt;223&gt; 12-646-433.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 439..458

&lt;223&gt; 12-646-433.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 6..26

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 474..494

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 426..450

&lt;223&gt; 12-646-433 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 31..32,35,59,514

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 371

ctacacttg	aaaatatctg	taccccaa	at	nnganagtca	atataacctg	gtgggtaana	60
gctggcactc	tggatccaga	tggccttgat	tcaagtctta	gaaaccactg	tataatgatg		120
tgggactgga	caagttactt	gaacttcatt	tgactcagct	tcctcaccaa	tgaaatggca		180
atagttgtca	tcacattatc	aatatcatgg	gttgctgaaa	gaacgaattc	attaatattg		240
caaagcacta	acaattgtgt	gtacagcaaa	ataaaattgt	actattatca	ttaaacaagt		300
caaaaaat	ccacaataat	aatttttagt	ttcatataat	tgcataatat	attatagttt		360
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tgtgggaatt	aaacttgkag	cataaaccaa	gggctgacaa	actatggcct	cttggctggg		480
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aaaattatac	actttaagca	tttaatcttt	atgattcatt	ggcatttaac	acatttgaaa		600
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ttgttgtgag	ttctctccaa	ttttatattg	aagtattttc	ttgagaatgg	ccataaacca		960
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&lt;210&gt; 372

&lt;211&gt; 1001

321

<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 503  
<223> 12-647-145 : polymorphic base A or G

<220>  
<221> misc\_binding  
<222> 483..502  
<223> 12-647-145.mis1, potential

<220>  
<221> misc\_binding  
<222> 504..523  
<223> 12-647-145.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 359..378  
<223> upstream amplification primer

<220>  
<221> primer\_bind  
<222> 788..808  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 491..515  
<223> 12-647-145 potential probe

<220>  
<221> misc\_feature  
<222> 351,951  
<223> n=a, g, c or t

<400> 372  
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tggtctgatt tctctgttat taaatatatt cagaggtggt ctgatttaca ttgttgtagag 180  
ttctctccaa ttttatattg aagtattttc ttgagaatgg ccataaacca agatcctgga 240  
aagctctgaa taaggtgggg aagtgagttt cagatatatg agtccggtaa aatataacga 300  
aatgagaaaag aaagaacaaa atgcaaaggt agaagaaatt tattactcac nagttcctaa 360  
gtgatgttag ggatgatgat aggaaactga cagaaagtcc agaggtggta gagagctcaa 420  
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gtattatttc ctaggcttct ccrcaggagt tgaggaatgg ctagcttaag ggaacacaca 540  
tgaaggggaa aattatcata tgactctggt attgatcatt aggttatatc atggatcatca 600  
cctctgtgat gtgttcggt tctgggtcat taggatgaga aacatgtaga ctctatctca 660  
aagaagggaa gttttaacaa aggcaatagt gacaacctat gactaggtct taagcaactc 720  
atgttaagcc taaaaatcaa tgctgaggca aaacaaactt acgacgagaa ggtagacaga 780  
cattttagaa gaaatgtgac ctagtgcgt ctggattggt agtataaaaa gtaccatgaa 840  
tcagcttcta aatgtaacaa gaataaatag tgaaaattgt ttttagagac aactatgcaa 900  
acatattagg agaagttgac tgcagtggca tgcacctata gttccagcta nctcaggagg 960  
ctgaggaagg tggattactt gagcccagga gttcaaggcc a 1001

<210> 373  
<211> 751  
<212> DNA  
<213> Homo Sapiens

<220>

322

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<221> allele
<222> 251
<223> 12-648-123 : polymorphic base A or G

<220>
<221> misc_binding
<222> 231..250
<223> 12-648-123.mis1, potential

<220>
<221> misc_binding
<222> 252..271
<223> 12-648-123.mis2, potential complement

<220>
<221> primer_bind
<222> 129..147
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 607..627
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 239..263
<223> 12-648-123 potential probe

<220>
<221> misc_feature
<222> 14,23,28,31..32,35,44,49,67,106,113,124
<223> n=a, g, c or t

<400> 373
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aatncaggaa aacaagtctt tagcaggagt tttctgaagc gtgaagtgag ctgctcaaga      180
gagtgaccgg atcttttctg gagaaaagcc tgggtggacct ctccactga agtgtgcatg      240
tgcactcctc rcataggaca tgggtgaggtg atgatgagag tccctcgaag ccagttgtac      300
ataacacatt ccatgagggt gccaatTTTA cagtaagata tcagagtaac aggtgtcacc      360
cataaacaaa ccagaaata tgtaactata catgtttact tatggattat cagtaggatt      420
ttgaggacga gttcacagaa attcaaaacc atacaccagt caaaaccatg caccaggctt      480
cgggatgtca gcaattctta ccttatgtga tgaaggagtg atttagaagc aaggtttggg      540
aaattctgac aaattttaag ataggttaga aataattata cagttgagag aaatatataa      600
tatcttccca atcataagtg acttttcctt agtagcataa tcaaggaaat gaaattataa      660
tgactacatt aaagtaatga tttagaaaac ctccctgaaa taaaaacaag ttacttagga      720
gaacaataac aatataaaat taaagagaag t                                     751

<210> 374
<211> 928
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 428
<223> 12-648-300 : polymorphic base C or T

<220>
<221> misc_binding
<222> 408..427
<223> 12-648-300.mis1, potential

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<220>
<221> misc_binding
<222> 429..447
<223> 12-648-300.mis2, complement

<220>
<221> primer_bind
<222> 129..147
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 607..627
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 416..440
<223> 12-648-300 potential probe

<220>
<221> misc_feature
<222> 14,23,28,31..32,35,44,49,67,106,113,124
<223> n=a, g, c or t

<400> 374
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aaattgnatt ttaaaacctc gtaagtagaa gggatcacaa gtaagnataa ttntcaaaca      120
aatncaggaa aacaagtctt tagcaggagt tttctgaagc gtgaagtgag ctgctcaaga      180
gagtgaccgg atcttttctg gagaaaagcc tgggtggacct ctccactga agtgtgcatg      240
tgcacacctc gcataggaca tgggtgagggt atgatgagag tccctcgaag ccagttgtac      300
ataacacatt ccattgagggt gccaatttta cagtaagata tcagagtaac aggtgtcacc      360
cataaacaata cccagaaata tgtaactata catgtttact tatggattat cagtaggatt      420
ttgaggayga gttcacagaa attcaaaacc atacaccagt caaaaccatg caccaggctt      480
cgggatgtca gcaattctta ccttatgtga tgaaggagtg atttagaagc aagggttggg      540
aaattctgac aaattttaag atagggttaga aataattata cagttgagag aaatatataa      600
tatcttccca atcataagtg acttttcctt agtagcataa tcaaggaaat gaaattataa      660
tgactacatt aaagtaatga tttagaaaac ctccctgaaa taaaacaag ttacttagga      720
gaacaataac aatataaaat taaagagaag taaacagtgg tagtccttct aagaatctat      780
attagtgatt acaagagctt ggttcccata tttttaactg tatctttccc tcactactgt      840
gtaaagatac taatttattt cttactttac atacttaatc aaatacatat acacacacac      900
gcacatgcac acacttacat aactacag                                     928

<210> 375
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-648-402 : polymorphic base G or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-648-402.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-648-402.mis2, potential complement

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<220>  
 <221> primer\_bind  
 <222> 100..118  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 578..598  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-648-402 potential probe  
  
 <220>  
 <221> misc\_feature  
 <222> 2..3,6,15,20,38,77,84,95,910..911,996..997  
 <223> n=a, g, c or t

<400> 375  
 gnnacntggg aattnctcgn ttacagaaa gaaattgnat tttaaaacct cgtaagtaga 60  
 agggatcaca agtaagnata attntcaaac aaatncagga aaacaagtct ttagcaggag 120  
 ttttctgaag cgtgaagtga gctgctcaag agagtgaacc gatcttttct ggagaaaagc 180  
 ctggtggacc tctccactg aagtgtgcat gtgcatcctc cgcataggac atggtgaggt 240  
 gatgatgaga gtccctcgaa gccagtgtga cataacacat tccatgaggg tgccaatttt 300  
 acagtaagat atcagagtaa caggtgtcac ccataaacia acccagaaat atgtaactat 360  
 acatgtttac ttatggatta tcagtaggat tttgaggacg agttcacaga aattcaaaac 420  
 catacaccag tcaaaacat gcaccaggct tcgggatgtc agcaattctt acctatgtg 480  
 atgaaggagt gatttagaag saaggtttgg gaaattctga caaattttta gataggttag 540  
 aaataattat acagttgaga gaaatatata atatcttccc aatcataagt gacttttcct 600  
 tagtagcata atcaaggaaa tgaaattata atgactacat taaagtaatg atttagaaaa 660  
 cctccctgaa ataaaaacia gttacttagg agaacaataa caatataaaa ttaaagagaa 720  
 gtaaacagtg gtagtcttcc taagaatcta tattagtgat tacaagagct tggttcccat 780  
 atttttaact gtatcttcc ctcactactg tgtaaagata ctaatttatt tcttacttta 840  
 catacttaat caaatacata cacacacaca cgcacatgca cacacttaca taactacaga 900  
 ttaataatan ngagaaaatt gatgttgtat aacataagtc tttccaaaagg aactgagcga 960  
 tgctgtattt aaaaaactgg cactactaga agtcanngtat t 1001

<210> 376  
 <211> 798  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 298  
 <223> 12-652-115 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 278..297  
 <223> 12-652-115.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 299..318  
 <223> 12-652-115.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 184..203



325

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 615..635

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 286..310

&lt;223&gt; 12-652-115 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 43..44,341

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 376

aaataattgg	ttaaaaattt	tatgccaca	aaattaaaca	ctnnaaccat	ttaatgtgta	60
caattcattg	gcgtttaata	tatttgcaat	attttgccac	taccaccact	ctctagatcc	120
aaagtatttt	gtcacttcaa	aaggaagctt	tatacccat	agacagtcac	ctcgcattct	180
cccatcctcc	atctcctgca	acctggcatt	atctgttggc	atcattctat	agcctagtag	240
ataatgtttt	tttaaaggca	aatagggtaca	taacttcaac	aaagtgtttt	atttttcytt	300
atttttacat	ctcctatttg	ttcctaaatg	aaaattctgt	ngatgtataa	gaattagtta	360
ttatttctta	gttgctctgt	tattaaatat	attcagcagt	gttggtgattt	atattgttac	420
agtttctgtc	caattttgta	ttgaagtctg	tccctttaga	attgcaataa	accaagctct	480
gatggagggtg	aggaagtga	attcagatgt	gtgtgtcagg	taaaatacaa	tgaaatgtaa	540
aataaaaacca	aaatgcatga	aatagaagaa	atgtattaca	gttcctagag	atattaagga	600
taatgacata	aaacggacag	aaaattcaga	agtggcagag	agctcaacca	gctgcctgga	660
agtgatagag	agaatccctg	tgtggggggc	tgtgggggtg	cttttcttaa	ggtctatgag	720
cattatttcc	tagtcttgtc	tgccggagtt	gtggagtggc	tatcttaaga	aaacacacac	780
gaagagggaa	aattatta					798

&lt;210&gt; 377

&lt;211&gt; 886

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 386

&lt;223&gt; 12-652-203 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 367..385

&lt;223&gt; 12-652-203.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 387..406

&lt;223&gt; 12-652-203.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 184..203

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 615..635

&lt;223&gt; downstream amplification primer, complement

326

```

<220>
<221> misc_binding
<222> 374..398
<223> 12-652-203 potential probe

<220>
<221> misc_feature
<222> 43..44,341
<223> n=a, g, c or t

<400> 377
aaataattgg ttaaaaattt ttatgccaca aaattaaaca cttnnaaccat ttaatgtgta      60
caattcattg gcgtttaata tatttgcaat attttgccac taccaccact ctctagatcc      120
aaagtatttt gtcacttcaa aaggaagctt tatacccatt agacagtcac ctgcgattct      180
cccatccctc atctcctgca acctggcatt atctgttggc atcattctat agcctagtag      240
ataatgtttt tttaaaggca aataggtaca taacttcaac aaagtgtttt atttttcttt      300
atttttacat ctctatttg ttctaaatg aaaattctgt ngatgtataa gaattagtta      360
ttatttccta gttgctctgt tattamatat attcagcagt gttgtgattt atattgttac      420
agtttctgtc caattttgta ttgaagtctg tccctttaga attgcaataa accaagctct      480
gatggagggtg aggaagtga aatcagatgt gtgtgtcagg taaaatacaa tgaaatgtaa      540
aataaaaacca aaatgcatga aatagaagaa atgtattaca gttcctagag atattaagga      600
taatgacata aaacggacag aaaattcaga agtggcagag agctcaacca gctgcctgga      660
agtgatagag agaatccctg tgtggggggc tgtgggggtg cttttcttaa ggtctatgag      720
cattatttcc tagtctgtc tgccggagtt gtggagtggc tatcttaaga aaacacacat      780
gaagaggggaa aattattatt tgattctggt gttgaccatt aggttatctc atggtaatca      840
gctctgtggt atgttgcatt tctgggtcag tgggatgaga aacaag                      886

<210> 378
<211> 957
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 457
<223> 12-652-274 : polymorphic base G or T

<220>
<221> misc_binding
<222> 437..456
<223> 12-652-274.mis1, potential

<220>
<221> misc_binding
<222> 458..477
<223> 12-652-274.mis2, potential complement

<220>
<221> primer_bind
<222> 184..203
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 615..635
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 445..469
<223> 12-652-274 potential probe

<220>

```

327

```

<221> misc_feature
<222> 43..44,341
<223> n=a, g, c or t

<400> 378
aaataattgg ttaaaaattt ttatgccaca aaattaaaca cttnnaaccat ttaatgtgta      60
caattcattg gcgtttaata tatttgcaat attttgccac taccaccact ctctagatcc      120
aaagtatttt gtcacttcaa aaggaagctt tataccatt agacagtcac ctgcattct      180
cccatectcc atctectgca acctggcatt atctgttggc atcattctat agcctagtag      240
ataatgtttt tttaaaggca aataggtaca taacttcaac aaagtgtttt atttttcttt      300
atttttacat ctctatttg ttctaaatg aaaattctgt ngatgtataa gaattagtta      360
ttatttccta gttgctctgt tattaaatat attcagcagt gttgtgattt atattgttac      420
agtttctgtc caattttgta ttgaagtctg tcccttkaga attgcaataa accaagctct      480
gatggagggtg aggaagtgaa attcagatgt gtgtgtcagg taaaatacaa tgaaatgtaa      540
aataaaacca aaatgcatga aatagaagaa atgtattaca gttcctagag atattaagga      600
taatgacata aaacggacag aaaattcaga agtggcagag agctcaacca gctgcctgga      660
agtgatagag agaatccctg tgtggggggc tgtgggggtg cttttcttaa ggtctatgag      720
cattatttcc tagtcttgtc tgccggagtt gtggagtggc tatcttaaga aaacacacat      780
gaagagggaa aattattatt tgattctggt gttgaccatt aggttatctc atggtaatca      840
gctctgtggt atgttgcat tctgggtcag tgggatgaga aacaagtgtg ctctgtaaca      900
aacaactaca caaggaaggg aagtttcaca aagccaaaag tgacaggcta tgacttg      957

<210> 379
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-652-371 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-652-371.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-652-371.mis2, potential complement

<220>
<221> primer_bind
<222> 131..150
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 562..582
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-652-371 potential probe

<220>
<221> misc_feature
<222> 288
<223> n=a, g, c or t

<400> 379

```

328

```

atgtgtacaa ttcattggcg ttaatatat ttgcaatatt ttgccactac caccactctc      60
tagatccaaa gtattttgtc acttcaaaaag gaagctttat acccattaga cagtcattctc    120
gcattctccc atcctccatc tcttgcaacc tggcattatc tgttggcatc attctatagc     180
ctagtagata atgttttttt aaaggcaaat aggtacataa cttcaacaaa gtgtttttatt    240
tttcttttatt tttacatctc ctatttggtc ctaaataaaa attctgtnga tgtataagaa    300
ttagttatta tttcctagtt gctctgttat taaatatatt cagcagtgtt gtgatttata    360
ttgttacagt ttctgtccaa ttttgatttg aagtctgtcc ctttagaatt gcaataaacc     420
aagctctgat ggaggtgagg aagtgaattt cagatgtgtg tgtcaggtaa aatacaatga     480
aatgtaaaat aaaacaaaaa ygcataaaat agaagaaatg tattacagtt cctagagata     540
ttaaggataa tgacataaaa cggacagaaa attcagaagt ggcagagagc tcaaccagct     600
gcctggaagt gatagagaga atccctgtgt ggggggctgt ggggggtgctt ttcttaaggt     660
ctatgagcat tatttcctag tcttgtctgc cggagttgtg gagtggctat ctttaagaaa     720
cacacatgaa gagggaaaat tattatttga ttctggtgtt gaccattagg ttatctcatg     780
gtaatcagct ctgtggtatg ttgcatttct gggtcagtgg gatgagaaac aagtgtactc     840
tgtaacaaac aactacacaa ggaagggaag ttccacaaag ccaaaagtga caggctatga     900
cttgctctta aacaactcaa gttaagtcta aaaatggatg tggaatcaat aactatattc     960
aacaatgta  tgacaagaag gtagaaaagg gatgggtcca g                                     1001

```

&lt;210&gt; 380

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-653-423 : polymorphic base T or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-653-423.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-653-423.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 903..921

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 390..410

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-653-423 potential probe

&lt;400&gt; 380

```

aaaaactgtg taaaaactgt tatctccaga gaaaaaccaa tgctaaggaa gcatccagtg      60
tagataatag agagtatcct ggagtcactg atattaataa tttagatgag agctgaacta    120
tatgcaggaa taggtaaaag aatgaagaag agaaaaaac acaaaaagaa aagcaggtaa     180
agtgttcagg acagttctca agactcaaaag ttagtttgc aagggaagata ctgagtaaga    240
atcagatgat gctgataggc aagataagag ccagatactc ctcaggagtt gaaatattta    300
ttaagcacat ttagggacta ctaaaaagag ttaagaaaag aaaaatgagg ataagattat     360
acttttttaa aaaagattcc aagatgttcg atggattaaa ttgtggaagg gccaaaactag    420
aaagagctga ccattgagga aattttgtat gaattcaggt agcagatgat ggaaaaatgg     480
actagaaagt ggatagaawt acctatgcct actttttacg taatatgact aacttcatat     540

```

329

```

tgtgttgtgt ggaaaaaagt taatacaaat aaaccactta aaatgtctct ggcataatagt 600
tagtgattca gaaatattat ttgcaatta tggttatttt tgttattact aatactatga 660
attacttaac atgtgtaagt cacttgagat attatccctc atttaataga aactaattat 720
tcagcatttc aagattatat tctctaacaa agtcctatgg tcttctatat gacagatacc 780
ctgtagactt gtttaaaata agacatatca gttttgacag taagatgagc aaaatgaaat 840
tgtaaaattc taccacaagt gacaaggatc ttcattcagta ttctaactta taagaaaata 900
atcccttacc gatataaatg tgtgattcta ttagtatttg cagccagacc cagtgttcac 960
ttgattttac taaagcattt aaatcattct gcgttgagat c 1001

```

```

<210> 381
<211> 1001
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> allele
<222> 499
<223> 12-654-115 : polymorphic base T or C

```

```

<220>
<221> misc_binding
<222> 479..498
<223> 12-654-115.mis1, potential

```

```

<220>
<221> misc_binding
<222> 500..519
<223> 12-654-115.mis2, potential complement

```

```

<220>
<221> primer_bind
<222> 595..613
<223> upstream amplification primer, complement

```

```

<220>
<221> primer_bind
<222> 76..96
<223> downstream amplification primer

```

```

<220>
<221> misc_binding
<222> 487..511
<223> 12-654-115 potential probe

```

```

<400> 381
ccatcaggaa gacccactac gttatctgag acaatggcaa aagctgacat atggtcttatt 60
cgaaactact gggattttca atttcctcac ccactcttac caaatgttga gttcgttgga 120
ggactccact gcaaacctgc caaaccccta ccgaaggtaa actattactg tttgttttgt 180
ctgctttgaa gtttcagtac gaatgggtct atattcattc aaagtgtttg acttacactg 240
gaagaaaggt ggaagtggga agagtaaagc agataccaat tagaaactga cgtacatggt 300
gatactatca caagtttatg aatttcatca ttattaccaaa taaagaggga tactaaagag 360
actttgaaaa tagggtttgt aaattaaagc tttgattatg caacatgtaa gaaggtagtg 420
gccatttcatt caaagaatat ttataaagag attagcacac accacaggta cgtgtatggg 480
acacagtttc tatcccaaya caccttacat tctattttga aagatagaat atatgcaagt 540
aataaaaact gtgtaaaaac tggtatctcc agagaaaaac caatgctaag gaagcatcca 600
gtgtagataa tagagagtat cctggagtca ctgatattaa taatttagat gagagctgaa 660
ctatatgcag gaataggtaa aagaatgaag aagagaaaaa aacacaaaaa gaaaagcagg 720
taaagtgttc aggacagttc tcaagactca aagtttagtt tgcaagggaag atactgagta 780
agaatcagat gatgctgata ggcaagataa gagccagata ctctcagga gttgaaatat 840
ttattaagca catttaggga ctactaaaaa gagttaagaa aagaaaatat gggataagat 900
tatacttttt aaaaaaagat tccaagatgt tcgatggatt aaattgtgga agggccaaac 960
tagaaagagc tgaccattga ggaaattttg tatgaattca g 1001

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330

<210> 382  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 499  
<223> 12-654-207 : polymorphic base G or A

<220>  
<221> misc\_binding  
<222> 479..498  
<223> 12-654-207.mis1, potential

<220>  
<221> misc\_binding  
<222> 500..519  
<223> 12-654-207.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 687..705  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 168..188  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 487..511  
<223> 12-654-207 potential probe

<400> 382  
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cttgtgtcat ctatcttttc tttttttttc ccccatcagg aagaccact acgttatctg 120  
agacaatggc aaaagctgac atatggctta ttcgaaacta ctgggatttt caatttcctc 180  
accactctt accaaatggt gagttcgttg gaggactcca ctgcaaacct gccaaacccc 240  
taccgaaggt aaactattac tgtttgttt gtctgcttg aagtttcagt acgaatggtt 300  
ctatattcat tcaaagtgtt tgacttacac tggaagaaag gtggaagtgg gaagagtaaa 360  
gcagatacca attagaaact gacgtacatg ttgatactat cacaagtta tgaatttcac 420  
cattattacc aataaagagg gatactaaag agactttgaa aatagggttg gtaaattaaa 480  
gctttgatta tgcaacatrt aagaaggtac tggccattca ttcaaagaat atttataaag 540  
agattagcac acaccacagg tacgtgtatg ggacacagtt tctatcccaa tacaccttac 600  
attctatttt gaaagataga atatatgcaa gtaataaaaa ctgtgtaaaa actgttatct 660  
ccagagaaaa accaatgcta aggaagcatc cagtgtagat aatagagagt atcctggagt 720  
cactgatatt aataatttag atgagagctg aactatatgc aggaataggt aaaagaatga 780  
agaagagaaa aaaacacaaa aagaaaagca ggtaaagtgt tcaggacagt tctcaagact 840  
caaagttagg tttgcaagga agatactgag taagaatcag atgatgctga taggcaagat 900  
aagagccaga tactcctcag gagttgaaat atttattaag cacatttagg gactactaaa 960  
aagagttaag aaaagaaaat atgggataag attatacttt t 1001

<210> 383  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 503  
<223> 12-657-396 : polymorphic base A or G

```

<220>
<221> misc_binding
<222> 483..502
<223> 12-657-396.misl, potential

<220>
<221> misc_binding
<222> 504..523
<223> 12-657-396.mis2, potential complement

<220>
<221> primer_bind
<222> 108..128
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 566..586
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 12-657-396 potential probe

<220>
<221> misc_feature
<222> 74,84,232,860,993
<223> n=a, g, c or t

<400> 383
ctgtataatt atcgggttgt gtgctgcttc tagtaaataa ttattatata ctgaagctat      60
atgtttctaa cacnaaactt tatnaatttc tgttctaaca tgcattctcag aatatagggga    120
caatatagct atatgtatga caacttcaaa attttatttt aaaagtatac ataaactata      180
ttattttatg tgggtattgca tctttataat gaagacaatt ttttctctgg angaaaatgg      240
aagaatttgc ccagagctct gatgaagacg gtggttgtgt ttctctggag tcagctgtgc      300
aaaaccttac agaagaaaaa gctgatctta tcacttcggc cctggctcag attccacaaa      360
aagtcagtac aacctccaat ccttataaga aactattcac acaatggaga aagtatggct      420
ttccacctgg aacttgaatc tcatttttca atttgcataa caggcactag atttatgtaa      480
caatttggaa agtattatgg tartttatgt gagcacaact gattatttgt ctagtgatct      540
ttgctattac tttagtaaca catctcttgg ttgtcgtttg ttaataataa agtaaaaata      600
aagcattaag tccctatttc acgttgcagg atttgaaatc ttaagaccta ttctgatgac      660
tccaaaggaa acttcttaag tatactagct caaaggaacc taaacttttg gggatgatat      720
aagaaagata gagaagaatc atgctcaata ttatcttcaa catattttat tgtataggag      780
ctctattcag tgtgtttgaa cataaaaagta gaagcttaga tttatgtagt ctttctaattg      840
aactggagtt ttctatagtn tacaaggcaa ttatgatttt aatattacag tctaacaacc      900
tgcattgtaat actttatgat attcaattaa ttttattact gtaattctag ttttcctctc      960
tttagttagt gctatttata cactagcctc aanggggttac t                               1001

<210> 384
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 12-658-120 : polymorphic base A or T

<220>
<221> misc_binding
<222> 483..502

```

332

&lt;223&gt; 12-658-120.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 12-658-120.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 384..404

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 863..883

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 12-658-120 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 418,551

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 384

```

atTTTTcaat ttgcataaca ggcactagat ttatgtaaca atttggaaag tattatggta      60
gtttatgtga gcacaactga ttatttgtct agtgatcttt gctattactt tagtaacaca      120
tctcttggtt gtcgtttggt aataataaag taaaaataaa gcattaagtc cctatttcac      180
gttgcaggat ttgaaatctt aagacctatt ctgatgactc caaaggaaac ttcttaagta      240
tactagctca aaggaacctt aactttgggg gatgatataa gaaagataga gaagaatcat      300
gctcaatatt atcttcaaca tattttattg tataggagct ctattcagtg tgtttgaaca      360
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caaggcaatt atgattttta tattacagtc taacaacctg catgtaatac tttatgatat      480
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tataggtaac ttctgtctct ttcttttttg attaaaaata aatgaaataa aggagagaat      840
gaagcataaa gagaaaagac aaccaactat ccaaaaccag acaaaagcca aagcaatgtt      900
tgtctctggg aaactgtaaa tttgataata gagctagaat ggccaagtga tttacattta      960
cacagcagtt gtgtgtccac aaggatattt aacttccata t                                1001

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&lt;210&gt; 385

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-659-382 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-659-382.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding



333

<222> 502..521  
 <223> 12-659-382.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 120..139  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 552..572  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-659-382 potential probe

<220>  
 <221> misc\_feature  
 <222> 21,546  
 <223> n=a, g, c or t

<400> 385  
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 tacattccact ttcaacttcc aaataatggt aagcttttca tttctaaagc aatgaaaagg 180  
 gagtgagttt acttatattc acttgtttca taagttatat tcccatatct gatgatttac 240  
 aaagaaagca aatctatcaa tcatctatgt ctgagaaata cattaatctg cttaaaaggc 300  
 acatataatt tgtttccgat taaaaattat taagcacttt ttccaagaaa aaaattcaag 360  
 tgactttttt tcccatatgt gcagaataat atcgcatgtg ctacaattgg aaattattta 420  
 aagggtggaaa agactgattg gcagttcttt aataataggt ctgtttaaaa tatcccaaat 480  
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 cgttgncagc agtgggtttat atgggtctgct atagctaagc catgtagtac atgcctagta 600  
 ccaacatgtg gtactgattt tggaattacc aagcaatctt cataaactag tgtataatct 660  
 acaagttttt aacgtgtact tgctatctga gttttaaaag taactacaca tttgcatcaa 720  
 aaaattagta taggtgatta caagaatggt atacaggata taaaactaaa aactgcttta 780  
 taaatattca aatatcaaag ttgctgcac agctcagaga cgctgaagct gtgctggagg 840  
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 agttttgcgt ccacatacct acctagggct attattaaca a 1001

<210> 386  
 <211> 983  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 306  
 <223> 12-660-134 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 286..305  
 <223> 12-660-134.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 307..326  
 <223> 12-660-134.mis2, potential complement

<220>

334

```

<221> primer_bind
<222> 173..193
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 692..712
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 294..318
<223> 12-660-134 potential probe

<220>
<221> misc_feature
<222> 880..881,889,925..926,930
<223> n=a, g, c or t

<400> 386
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ataacagccc ttcttttatt accagtgaca gcacttttta tgggaataaa tccccagtct      180
ttattatgaa gtggtaacac attttgtcat ggagtgtggc ctgtccttct actactggag      240
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tcactrtttc tctaagagaa tatgttctag catgcacctc gctttttggg ggctagagtg      360
ggggatatgc agtttttttt gtctgcagtg tgctctttct atttgtccac ttgacaatct      420
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tcgccccaga ggacttggtg cttctttctc tgagatctga aagtaatttt aatgcagttt      540
tactgttaca gtaatttggt acctagagaa ccattgcccatttcattgcct catattgagc      600
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ccatgggata cctatgggtg gccttccttt gttttagatg caaccgata agttggttca      780
catgaaggct aagggggaag ctgttacagt agactcaaaa acgatgtcat ctacaaattt      840
gctcaatgca taaaaagcag tcattaatga ctcttcatgn ntgtaattnt tttaactacg      900
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ccactatgag tatcacgata tgt                                     983

<210> 387
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 497
<223> 12-662-80 : polymorphic base G or C

<220>
<221> misc_binding
<222> 477..496
<223> 12-662-80.misl, potential

<220>
<221> misc_binding
<222> 498..517
<223> 12-662-80.mis2, potential complement

<220>
<221> primer_bind
<222> 418..435
<223> upstream amplification primer

```

335

<220>  
 <221> primer\_bind  
 <222> 979..1000  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 485..509  
 <223> 12-662-80 potential probe

<220>  
 <221> misc\_feature  
 <222> 24,726,739  
 <223> n=a, g, c or t

<400> 387  
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 cagattattc agcacatcaa gggttatattg tcttggaag tcatagatga cagataccct 180  
 tggacttgat taaaagtaga catatcagtt gtgacagcaa gataagctag ttgaaatttt 240  
 aaaattcttc cataagtaat aaggatcttc accagtattc cagcttaaaa cacttcctca 300  
 acaatataaa tgtgtgcctc aaatatgtgc agacaaacac aggggttcaca tgattttaat 360  
 aaagcattta aatcattctg cattgagatc ccagaatttt acattttaga gcataaacat 420  
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 gaactcgtct tgacatsata ggtgtctgac agagaagcac agagataatg aacaatgcat 540  
 gtataataaa gaccaaataa ttcttgccac ttgtttctga atagtgccct tagtttcaat 600  
 acgaaaaaaa attcgcgtcag catataagat tatattcttt tcgaaggaat acaagaactt 660  
 catatatattt aattaacatc acattttaag gcatatgtaa aaagtaaggc ttcttcaatg 720  
 gattanttat gcaatctcnt taaaagatca tttttgcttg cataaaactg agaatttggt 780  
 ccattgttaa aactcagtat ctatgtttta tgcaagttgt ataggcttta tagagtcagt 840  
 ttcttaagag acaaaagtgt aggttaagact aatgaaaaat atataccact ctaccctact 900  
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<210> 388  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-906-149 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-906-149.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-906-149.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 353..372  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 809..829  
 <223> downstream amplification primer, complement

```

<220>
<221> misc_binding
<222> 489..513
<223> 12-906-149 potential probe

<220>
<221> misc_feature
<222> 750,853..854,860,942,945
<223> n=a, g, c or t

<400> 388
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agaaggcttc ctgtagtcaa ttaacaaaga ctttgagtga gtcaatactg ttacctcatg      180
gccctcact atgatctcct ctagaatgac cttgacatta agccaatggc tcatgtcaca      240
gggccacacc aggaccttct cacagaattc acaactagca cagaagagtt gcaggagccg      300
aaataccaaa gctgacttct cgggcctcat gatgacattt ccctcacaca ctgatctgca      360
ctagctttgt agttactaag catgaaattg aaatgacaat ataagcacag aagttaaaaa      420
ttaatatattt aagtgtaaat aaagttcatt gggtggttgc cagcttcaca tttattgaag      480
atatcaaaaa gaaatgcaat rttgcaatgt atcagggaaa atttggtcca accttctaga      540
gcttagaaat aaaacacata aagatacttt tattatctgg attgatgata aagaaatttt      600
ttaaaattct ttagatttaa taattcatgt gcagaaacat atgcacacaa ccacatttac      660
attcattcca cactaaatcc aaggtgtcag atggtttaga agaactcatg tttcacatcc      720
tttgctcaca aggacaatac aaagaattan ctgggaatac acttggagac tttgggtaac      780
tattttcatt aattctatat actaagtgc aagtgttcag tatttcagag gagaaaaatg      840
cacaattgct ttnnaatatn ctgataatta ataaaagctt tagtaagaca tttgacttag      900
gaggtgagat gaaagcttca ttaggattta ttgttgccaa anggnaagca tttcacaaac      960
aatacaagca tgacgaaagt ataggatatt taaaagtcac t                        1001

<210> 389
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-906-154 : polymorphic base A or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-906-154.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-906-154.mis2, potential complement

<220>
<221> primer_bind
<222> 348..367
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 804..824
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513

```

337

&lt;223&gt; 12-906-154 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 745,848..849,855,937,940

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 389

ttataactga	ttgccagggt	gttaagcctg	gcaagacatt	cagagctagg	tcaacaaata	60
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gcttcctgta	gtcaattaac	aaagactttg	agtgagtcaa	tactgttacc	tcattggcccc	180
tcactatgat	ctcctctaga	atgaccttga	cattaagcca	atggctcatg	tcacagggcc	240
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ccaaagctga	cttctcgggc	ctcatgatga	catttccctc	acacactgat	ctgcactagc	360
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aaaaagaaat	gcaatgttgc	matgtatcag	ggaaaatttg	ttccaacctt	ctagagctta	540
gaaataaaac	acataaagat	acttttatta	tctggattga	tgataaagaa	attttttaaa	600
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ttccacacta	aatccaaggt	gtcagatggt	ttagaagaac	tcattgtttca	catcctttgc	720
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ttgcttttna	atatnctgat	aattaataaa	agcttttagta	agacatttga	cttaggaggt	900
gagatgaaag	cttcattagg	atattattgt	gccaaanggn	aagcatttca	caatcaatac	960
aagcatgacg	aaagtatagg	atatttaaaa	gtcattccag	t		1001

&lt;210&gt; 390

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-906-251 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-906-251.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-906-251.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 251..270

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 707..727

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-906-251 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

338

&lt;222&gt; 648,751..752,758,840,843

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 390

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cacctcaaat ttcagtgcag aaggcttccct gtagtcaatt aacaaagact ttgagtgagt      60
caatactggt acctcatggc ccctcactat gatctcctct agaatgacct tgacattaag      120
ccaatggctc atgtcacagg gccacaccag gaccttctca cagaattcac aactagcaca      180
gaagagttgc aggagccgaa ataccaaagc tgacttctcg ggcctcatga tgacatttcc      240
ctcacacact gatctgcact agctttgtag ttactaagca tgaaattgaa atgacaatat      300
aagcacagaa gttaaaaatt aatattttta gtgtaaataa agttcattgg ttggttgcca      360
gcttcacatt tattgaagat atcaaaaaga aatgcaatgt tgcaatgtat caggggaaat      420
ttgttccaac cttctagagc ttagaaataa aacacataaa gatactttta ttatctggat      480
tgatgataaa gaaatttttt waaattcttt agatttaata attcatgtgc agaaacatat      540
gcacacaacc acatttacat tcattccaca ctaaatccaa ggtgtcagat ggtttagaag      600
aactcatggt tcacatcctt tgctcacaag gacaatacaa agaattanct gggaatacac      660
ttggagactt tgggtaacta ttttcattaa ttctatatac taagtgcaaa gtgttcagta      720
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gtaagacatt tgacttagga ggtgagatga aagcttcatt aggatttatt gttgccaaan      840
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cagtgagtag tgatcacaga gaaagttcag ggcaggacgt tacagaacaa agttagtgtg      960
aaataggaga aattagtttt aaataaacia ttattagtct g                               1001

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&lt;210&gt; 391

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-906-451 : polymorphic base A or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-906-451.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-906-451.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 52..71

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 508..528

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-906-451 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 449,552..553,559,641,644

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 391

339

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aataccaaaag ctgacttctc gggcctcatg atgacatttc cctcacacac tgatctgcac      60
tagcttttga gttactaagc atgaaattga aatgacaata taagcacaga agttaaaaaat    120
taatatttta agtgtaaata aagttcattg gttgggtgcc agcttcacat ttattgaaga    180
tatcaaaaag aaatgcaatg ttgcaatgta tcagggaaaaa tttgttccaa ccttctagag    240
cttagaaaata aaacacataa agatactttt attatctgga ttgatgataa agaaattttt    300
taaaattctt tagatttaat aattcatgtg cagaaacata tgcacacaac cacatttaca    360
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ttgctcaca ggacaataca aagaattanc tgggaataca cttggagact ttgggtaact    480
attttcatta attctatata mtaagtgcaa agtggtcagt atttcagagg agaaaaatgc    540
acaattgctt tnnaatatnc tgataattaa taaaagcttt agtaagacat ttgacttagg    600
aggtgagatg aaagcttcat taggatttat tgttgccaaa nggnaagcat ttcacaatca    660
atacaagcat gacgaaagta taggatattt aaaagtcatt ccagtgaagta gtgatcacag    720
agaaagtcca gggcaggacg ttacagaaca aagttagttt gaaataggag aaattagttt    780
taaataaaca attattagtc tgaatgataa tttataggta aataattact tactcaagtt    840
ttaaaggaca gtcacaagat cgagcttttc cttagcaaga aattcttgca tctcaacata    900
gagtgatatg taaagataag agtttagaaa cagggaaaat gataaaaatt gtatattagt    960
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&lt;210&gt; 392

&lt;211&gt; 744

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 244

&lt;223&gt; 12-907-199 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 224..243

&lt;223&gt; 12-907-199.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 245..263

&lt;223&gt; 12-907-199.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 46..65

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 533..553

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 232..256

&lt;223&gt; 12-907-199 potential probe

&lt;400&gt; 392

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tgacttttagg aacatttaga aagtttaaag taaacaatgc tcaatgtaat tgagcatgag      60
gagtgggtga attatgattt aaaaccagcc ttgtcttttt ttatagtgac ttgaagagggt    120
ttagtaattc cttcatgttt tggaccgaga ccgattccag gaacaaacgt catgtttttcc    180
attatatata gactgggaac actataagag ctatgtggaa tattaattgc agccccccat    240
tgtkccagct aatctctgcc tcaaagatta atggggattg gtgtgatata aggctgaatt    300
gtaccttttg accatcaggg ccagtgcaag gcaagataaa tgtgctctgg tgaacttcat    360
cagctttttc aaccctact tatcccatgt tagtgggatg tttaagccag aaggaaggcc    420
atacattaga ggaaataata gaaacatcag cccagcatc cactatgccc tcaaaccctt    480
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340

cggttttttc actgttggag cctatcccag agccccctgt cttatctcct ttgttttaaaa	600
caatattagg taataaaagt aattgagcaa ttgactcacc aactggaatg gaaacaggaa	660
ccttggcaga caccataagt ttaatctcat cagaggaatc agaattaatg agaccagtag	720
gaactatgat accttttagca gagg	744

&lt;210&gt; 393

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-907-482 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-907-482.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-907-482.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 20..39

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 507..527

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-907-482 potential probe

&lt;400&gt; 393

aaagtaaaca atgctcaatg taattgagca tgaggagtgg ttgaattatg atttaaaacc	60
agccttgtct ttttttatag tgacttgaag aggttttagta attccttcat gttttggacc	120
gagaccgatt ccaggaacaa acgtcatgtt ttccattata tatagactgg gaacactata	180
agagctatgt ggaatattaa ttgcagcccc ccattgttcc agctaattctc tgcctcaaag	240
attaatgggg attggtgtga tataaggctg aattgtacct tttgaccatc agggccagtg	300
caaggcaaga taaatgtgct ctggtgaact tcatcagctt ttccaacccc tacttatccc	360
atgttagtgg gatgtttaag ccagaaggaa ggccatacat tagaggaaat aatagaaaca	420
tcagccccag catccactat gccctcaaac ctttttcctt gaatgtgtat ggtgcagggtg	480
ggccattgtt tagaaattac wttaatccaa taagcggctt tttcactgtt ggagcctatc	540
ccagagcccc ctgtcttctc tcctttgttt aaaacaatat taggtaataa aagtaattga	600
gcaattgact caccaactgg aatggaaaca ggaaccttgg cagacaccat aagtttaatc	660
tcatacaggg aatcagaatt aatgagacca gtaggaaacta tgataccttt agcagagggtg	720
gatgtcctac ctaacaccag gcccatggaa ccttgaggta aagggccagt gacccctgtg	780
gggacaatta aaagacaaga attaggtagt aaatttagag gaatggtact atggagatca	840
accaccctgc cccctactgt ggaggtagac aagcagtgtg ctgtgacaga agaagaggct	900
gggacccatc tgggtttgct gtaggtaaat ttgtttgtac tgggggctgc gttaggaccg	960
cttcaattgg aaatgcaaca ttggtctgag tctgagggtgt g	1001

&lt;210&gt; 394

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens



341

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<220>
<221> allele
<222> 53
<223> 12-909-36 : polymorphic base A or G

<220>
<221> misc_binding
<222> 33..52
<223> 12-909-36.mis1, potential

<220>
<221> misc_binding
<222> 54..73
<223> 12-909-36.mis2, potential complement

<220>
<221> primer_bind
<222> 18..38
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 505..525
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 41..65
<223> 12-909-36 potential probe

<220>
<221> misc_feature
<222> 480,526,538..539,545,554..556,573
<223> n=a, g, c or t

<400> 394
taacttgtgt tttgactcca ggaaaagtaa aatgttagtt tgagctcgag aargaatctg      60
gatactggta actttaccca gaccttgga atcttcccc tcagtacatt taattaatga      120
agtgttacaa tgaattctca aaagatctaa gagatttggt ttcacttcat tcgctgtgat      180
catgggccta atcagagtca ctgcactggc aaccactgct ggagtggcat tataccaatc      240
tattcaaaca gctcattttg tcaatgatta gcaagtcaat tccacccaaa tgtggaattc      300
tcaacaagac attgatcaaa aattagctaa tcaaattaat gatttaagac agtctgttat      360
ttggcttgga aatcagctga tgagtctcga acatcacatg caaatgcagt gtgatttgaa      420
tacttctgat ttctgtatca caccatattc ctacaatgag actgatcatt catggaaaan      480
tgggtcaaagg acaccttctg ggtagggaag ataatttatt cttggnacat aactaaannt      540
taaangaaac aaannntttc tgaagcctct cangctcatt taccatttgt gtctggagct      600
gaggcgtag atcagggtggc agaaagtatt tctggactaa accccacgac ttggattaag      660
tctactgggg gctccatggt agtaaatttt ggaataatat ttctctgttt aatcgacttg      720
tttttagtgt gctggaccag tcaaagattc ccgtgtcaaa acctagagaa caaacaagac      780
tttgcaccat ggcacattta tataaaaaga aagggaggga tgttgtggga agtcaggggac      840
cctgaatgga gggactggct ggagctgttg cagaggaaca taaattgtga agatttcatt      900
ttaatatgga cctttgtcag ttcccaaata atacttttct aatttcttat gcctgtctta      960
ctttaatctc ttaatcctgt tatcttcata acctgaggat g                                1001

<210> 395
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 193

```

342

&lt;223&gt; 12-909-176 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 173..192

&lt;223&gt; 12-909-176.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 194..213

&lt;223&gt; 12-909-176.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 18..38

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 505..525

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 181..205

&lt;223&gt; 12-909-176 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 480,526,538..539,545,554..556,573

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 395

taacttggtg	tttgactcca	ggaaaagtaa	aatgtaggtt	tgagctcgag	aaggaatctg	60
gatactggta	actttacc	gaacctggga	atcttcccc	tcagtacatt	taattaatga	120
agtgttacaa	tgaattctca	aaagatctaa	gagatttgtt	ttcacttcat	tcgctgtgat	180
catgggccta	atyagagtca	ctgcactggc	aaccactgct	ggagtggcat	tataccaatc	240
tattcaaaca	gctcattttg	tcaatgatta	gcaagtcaat	tccacccaaa	tgtggaattc	300
tcaacaagac	attgatcaaa	aattagctaa	tcaaattaat	gatttaagac	agtctgttat	360
ttggcttga	aatcagctga	tgagtctcga	acatcacatg	caaatgcagt	gtgatttgaa	420
tacttctgat	ttctgtatca	caccatattc	ctacaatgag	actgatcatt	catggaaaan	480
tggtcaaagg	acaccttctg	ggtagggaag	ataatttatt	cttggnacat	aactaaannt	540
taaaangaaac	aaannntttc	tgaagcctct	cangctcatt	taccattgt	gtctggagct	600
gaggcgtag	atcaggtggc	agaaagtatt	tctggactaa	acccacgac	ttggattaag	660
tctactgggg	gctccatggg	agtaaatttt	ggaataatat	ttctctgttt	aatcgacttg	720
tttttagtgt	gctggaccag	tcaaagattc	ccgtgtcaaa	acctagagaa	caaacaagac	780
tttgcaccat	ggcacattta	tataaaaaga	aaggagggga	tggtgtggga	agtcagggac	840
cctgaatgga	gggactggct	ggagctgtgg	cagaggaaca	taaattgtga	agatttcatt	900
ttaatatgga	cctttgtcag	ttcccaaata	atacttttct	aatttcttat	gcctgtctta	960
ctttaatctc	ttaatctgt	tatcttcata	acctgaggat	g		1001

&lt;210&gt; 396

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-909-484 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

343

<222> 481..500  
 <223> 12-909-484.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-909-484.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 18..38  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 505..525  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-909-484 potential probe

<220>  
 <221> misc\_feature  
 <222> 480,526,538..539,545,554..556,573  
 <223> n=a, g, c or t

<400> 396  
 taacttggtggt tttgactcca ggaaaagtaa aatgttagtt tgagctcgag aaggaatctg 60  
 gatactggta actttaccca gaccttggga atcttccccc tcagtacatt taattaatga 120  
 agtggttaciaa tgaattctca aaagatctaa gagatttggt ttcacttcat tcgctgtgat 180  
 catgggccta atcagagtca ctgcactggc aaccactgct ggagtggcat tataccaatc 240  
 tattcaaaca gctcattttg tcaatgatta gcaagtcaat tccacccaaa tgtggaattc 300  
 tcaacaagac attgatcaaa aattagctaa tcaaattaat gatttaagac agtctgttat 360  
 ttggcttgga aatcagctga tgagtctcga acatcacatg caaatgcagt gtgatttgaa 420  
 tacttctgat ttctgtatca caccatattc ctacaatgag actgatcatt catggaaaaa 480  
 tgggtcaaaagg acaccttctg kgtagggaag ataatttatt ctgggnacat aactaaannt 540  
 taaangaaac aaannnttct tgaagcctct cangctcatt taccattgt gtctggagct 600  
 gaggcgttag atcaggtggc agaaagtatt tctggactaa accccacgac ttggattaag 660  
 tctactgggg gctccatggt agtaaatttt ggaataatat ttctctgttt aatcgacttg 720  
 tttttagtgt gctggaccag tcaaagattc ccgtgtcaaa acctagagaa caaacaagac 780  
 ttgacacat ggcacattta tataaaaaga aaggaggagg tggtgtggga agtcaggggac 840  
 cctgaatgga gggactggct ggagctgtgg cagagggaaca taaattgtga agatttcatt 900  
 ttaatatgga cctttgtcag ttcccaaata atacttttct aatttcttat gcctgtctta 960  
 ctttaattctc ttaatcctgt tatcttcata acctgaggat g 1001

<210> 397  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 347  
 <223> 12-910-76 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 327..346  
 <223> 12-910-76.mis1, potential

<220>

344

<221> misc\_binding  
 <222> 348..366  
 <223> 12-910-76.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 272..292  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 704..724  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 335..359  
 <223> 12-910-76 potential probe

<220>  
 <221> misc\_feature  
 <222> 17,38,54,57,59,62,69,76,87,95,106,109,115,118,122,128,857,876,888  
 <223> n=a, g, c or t

<400> 397  
 aataattata cagaacnttt aagtgagaaa tgtccagntt gtttaatat tggagnent 60  
 cnaattctna aaacanaatg gtttgcnaac tcatnaaact atttcnacna tatcnagngt 120  
 cnatttgntg acattttgtg atgtcaacaa aatgataaaa ttttagaaac ccaacatcaa 180  
 ttccctcaca aaagtacaac cagtagctat ccaaatacaa aaatgccact ctgaattcac 240  
 cagatctcaa ggcagaagga gaaaacccga agactcacag acatgagaaa acttatgatt 300  
 ggtaagatga ttaatttttt tggactgtac caccocgctt ccacaarcca aaatgacacc 360  
 actgtgaggg aacttccttt cacctgcagt tattgaaatg ggaggaggga atttcaagtg 420  
 gacatttaat ttcttcattg atctgtaaatt tgaggggaa agcccacatt tgtcccaccc 480  
 cacagacggt attaagagt ttcagagggc tgaaccacct ggggtaaatt ggggacagga 540  
 gatggagtag taatcacagt gatcaccaca cagatcttgg catctgcttt gtgttcctag 600  
 atacggggat gccacacaga ggagtctcac cagaaccata gactgcagg gggcaaaatc 660  
 ttaaggaagg cctgaatctt tgacaaaatt ttacaattcc caggtagtca tgtggagatt 720  
 ttccatgact gggaaacaag tataagactt gaaattaagt tccaaggctt gtttaaattg 780  
 ctcccagatc tagaaatcac tgcaagtctg ggtttaagta ccagtgcagc atttaagttc 840  
 agatgctcac aataagntct cccaagactg gaaaanaca attaaagnaa atactgagtt 900  
 ttggtgcagt attaaattct ggtggcaaat attcagtcct tgcctaaaca aaaagcaact 960  
 ggtggcaagg aattagattc caatattaag tagtaaaggt t 1001

<210> 398  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 12-910-295 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 12-910-295.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..523  
 <223> 12-910-295.mis2, potential complement

345

```

<220>
<221> primer_bind
<222> 209..229
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 641..661
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 12-910-295 potential probe

<220>
<221> misc_feature
<222> 6,13,24,32,43,46,52,55,59,65,794,813,825
<223> n=a, g, c or t

<400> 398
attctnaaaaa canaatgggtt tgcnaactca tnaaactatt tcnacnatat cnagngtcna      60
tttgntgaca ttttgtgatg tcaacaaaat gataaaattt tagaaaccca acatcaattc      120
cctcacaaaa gtacaaccag tagctatcca aatacaaaaa tgccactctg aattcaccag      180
atctcaaggc agaaggagaa aaccggaaga ctcacagaca tgagaaaact tatgattggg      240
aagatgatta attttttttg actgtaccac cccgcttcca caagccaaaa tgacaccact      300
gtgaggggaa ttccctttcac ctgcagttat tgaaatggga ggaggggaatt tcaagtggac      360
atttaatttc ttcattggatc tgtaaatttg aggggaaagc ccacatttgt cccacccac      420
agacggtatt aagagtgttc agagggtga accacctggg gttaaattggg gacaggagat      480
ggagtagtaa tcacagtgat caycacacag atcttggcat ctgctttgtg ttcctagata      540
cggggatgcc acacagagga gtctcaccag aaccatagca ctgcaggggg caaaatctta      600
aggaaggcct gaatctttga caaaatttta caattcccag gtagtcatgt ggagattttc      660
catgactggg aaacaagtat aagacttgaa attaagttcc aaggcttgtt taaatgtctc      720
ccagatctag aaatcactgc aagtctgggt ttaagtacca gtgcagcatt taagttcaga      780
tgctcacaat aagntctccc aagactggaa aancaacatt aaagnaaata ctgagttttg      840
gtgcagtatt aaattctggg ggcaaatatt cagtccttgc ctaaacaaaa agcaactggg      900
ggcaaggaaat tagattccaa tattaagtag taaaggttga acaacacaag aatacaccta      960
taaaagttag aacatgtgga tatctcttta agtatggaaa c                                1001

<210> 399
<211> 740
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 240
<223> 12-911-22 : polymorphic base G or C

<220>
<221> misc_binding
<222> 221..239
<223> 12-911-22.mis1

<220>
<221> misc_binding
<222> 241..260
<223> 12-911-22.mis2, potential complement

<220>
<221> primer_bind
<222> 219..237
<223> upstream amplification primer

```

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<220>
<221> primer_bind
<222> 653..673
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 228..252
<223> 12-911-22 potential probe

<220>
<221> misc_feature
<222> 354,703,732,739
<223> n=a, g, c or t

<400> 399
atattttcca ctgtaatatt ttatcttagc tatatagcct aaaattttaa caattaatct      60
cggggggggag tgtcgcattc ttttgttgta ttttcctagg cgatttgatc cttttctaat      120
acattgctga cacaagatca ctaacaattc taaaatgtcc tattcatgat atcactatct      180
ttatgatgc acatttttta aaccttagtt gatatgttta ttcacaatat gctggtccts      240
tgtagtgcat aaatatgaac tgacatcaat agatgtaaaag aagtcaaaag cataaaattc      300
acaggaggca atagactata caagtcattg aagtggttcta tgatattagt ttgntattaa      360
aaacataaca atgttcctt tttgataaaa ataagccatt ttacatagcc aacagtactg      420
ggcttagacc atgaattggt aatgataata atgatagtgc attactctgg aaaagggtta      480
gtatatttct ctagaactca tctagatatt atggcctaca tttctgcctt tagtccaata      540
ttttttgtgt cctcaatgag aaagagatag ttatcatcaa tgtgttcaaa caaaagggtta      600
agtgatcctt tcttagggag agaatatgtc aacaagggtga tcaagttgaa cagattattt      660
taggagaggt aacgacagga aacattttat taaaaatact aantttttaa aaactttgat      720
cgtttacttg ancaaagant                                     740

<210> 400
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-912-65 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-912-65.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-912-65.mis2, potential complement

<220>
<221> primer_bind
<222> 437..457
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 908..928
<223> downstream amplification primer, complement

<220>
<221> misc_binding

```

347

<222> 489..513  
 <223> 12-912-65 potential probe

<220>  
 <221> misc\_feature  
 <222> 24,47,131,137,440  
 <223> n=a, g, c or t

<400> 400  
 acaacaggaa gctgcaccct tctnctgcat tcagacagaa gtgaganctg ctgggccgga 60  
 agctctagca aggatgatag caatacaagt tgattccttaa ttaattaaag cacattgaga 120  
 agctgatcca naaaagncaa acactattaa atctctcaga ctttaaaaat acagtcatat 180  
 tccaggagtt aaggggtcca gtcaccaac atggcacatg tatacatacg taacaaacct 240  
 gtacgttggtg cacatgtacc ctagaactta aagtataata aaaaatatat atatacatag 300  
 tcatattcat ttttcacaac atagcatatt gtgacaggtc tttcttgatt tactaagttt 360  
 acttattatc ctgtgaagta tattttcata cctgaatttt acctagcttc agccttccat 420  
 taacgttgca ttcattcttn aacatttgta aaccaccata ttttctagaa attggtgact 480  
 atattctaaa atcattgtag ycacgaaatc atatgaatac ctgcttgcc tctccggata 540  
 ggggtcaagtt tctcacataa gcaaaattta cttatttggc atatacaatg tgaacataat 600  
 cttaaaacct gactttggct tttggtgta ttaataaggt tcttaatcca atttttgttt 660  
 tttttttaca gaagtgtac agctatactg taatcctagc attcctgagt tttgtatata 720  
 gcttggaat aatattttatt gtcaaattt tttaaatct tcaacttctt ctaatatatt 780  
 tattaacctt agcatctttt tttgcatttt ttccctgtga agtgtcatte actctatcaa 840  
 ttactatcag aaaaatctgt tatcatcttg cctatttatg caaagcaaat gattctccaa 900  
 ctgttgactc taatccttcc tctcaaatag gttctgctaa ataaatgacc ctgagtctct 960  
 gagacatttg ttgtatcaaa gcacaacaac attaacagta a 1001

<210> 401  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 384  
 <223> 12-914-106 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 365..383  
 <223> 12-914-106.mis1

<220>  
 <221> misc\_binding  
 <222> 385..404  
 <223> 12-914-106.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 279..298  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 773..793  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 372..396  
 <223> 12-914-106 potential probe

<220>

348

<221> misc\_feature  
 <222> 536,990  
 <223> n=a, g, c or t

<400> 401  
 aactcctatt ataggaggt agttttctta ataatgttg atactttcca tatgcctact 60  
 tcactaattt taattatttc tctttttttg ctcttctgat tgggtaatta catatgtttt 120  
 atcttttagga ttgctgattc cttactctgt ttgaccaagt ctgctgttga agttttctac 180  
 tgagtttttc aattcacttc acataatttt tattaattgg atttcttttt atttttaaaa 240  
 cattttccact tctttgcaaa atttatcatc attttcctgg attattctct aaagtgtggt 300  
 ttttttcaac tctctattca tatgtgctgt gattttctgaa ctttttaaaat aggtgtgcac 360  
 tgaatttctt gtcaaaaatt ttayggagca tctgaacctt aattctgcac tgtaacttaa 420  
 aactagactt gcagtgattt ctaggctctg gagatactta agcaataact ggagcttaat 480  
 ctcaaagtgt atacttggtt gcaggtcata gaaaagctcc gtatgagtaa ctgggnatta 540  
 taaaaatacc tgccaaagat tcaggccttc ctttggtatt tggccctgc agtggtatgg 600  
 aactggtcag actcctcagc gtggcattcc tgctaatagg aacaccaagc agttgctgag 660  
 atgtgtgtgc ctgtcactgt gatttagtact tccactcttg tttccaattc accccaggtc 720  
 attcaactct atagccactc ctaatgcctc tcctgaggta ggacaaaagt gggcttccca 780  
 aacaaaaaag attcacagat ccatgaagaa actgaacatt cacttcaaatt ttcctcctct 840  
 gatcttgcca actgcaggta aaataaagt ctgtcacagt ggtgtcttct tgggtgtgtg 900  
 aaggggtgcc ataatccaaa aaattattca ttttacagat catgagtttt cccatgtcta 960  
 tgagtcttgg gattttcttc ttctcttttn agctctggtg a 1001

<210> 402  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 12-914-252 : polymorphic base A or T

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 12-914-252.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..523  
 <223> 12-914-252.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 252..271  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 746..766  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 12-914-252 potential probe

<220>  
 <221> misc\_feature  
 <222> 509,963  
 <223> n=a, g, c or t



349

```

<400> 402
ttaataatgt ttgatacttt ccatatgcct acttcactaa ttttaattat ttctcttttt 60
ttgctcttct gattgggtaa ttacatatgt tttatcttta ggattgctga ttccttactc 120
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tcattttaca gatcatgagt tttcccatgt ctatgagtct tgggattttc ttcttctctt 960
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<210> 403
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 10-448-266 : polymorphic base A or C

<220>
<221> misc_binding
<222> 483..502
<223> 10-448-266.mis1, potential

<220>
<221> misc_binding
<222> 504..523
<223> 10-448-266.mis2, potential complement

<220>
<221> primer_bind
<222> 238..257
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 660..679
<223> downstream amplification primer, complement

<220>
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<222> 491..515
<223> 10-448-266 potential probe

<220>
<221> misc_feature
<222> 772,907,915
<223> n=a, g, c or t

<400> 403
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agtttttact ttagctctgg gagttgtgga aagggtctgg tatgggccgc agaatacagc 120
ctttggatga atatgaagac aatcctgaaa gaacttgttc agagagggtca tgaggtgact 180

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350

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gtactggcat cttcagcttc cattcttttt gatcccaacg actcatccac tcttaaactt 240
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aagagattgt cagaaattca aaaagataca ttttggttac ctttttcaca agaacaagaa 360
atcctgtggg caattaatga cataattaga aacttctgta aagatgtagt ttcaaataag 420
aaacttatga aaaaactaca agagtcaaga ttgacatcg tttttgcaga tgcttattta 480
ccctgtgggt agctgctggc tgmgtatatt aacataccct ttgtgtacag tcacagcttc 540
agtcttggt actcatttga aaggcacagt ggaggattta tttccctcc ttcctacgta 600
cctgttggt tgtcaaaatt aagtgatcaa atgactttca tggagagggg aaaaaatatg 660
ctctatgtgc tttattttga cttttggttc caaatattta atatgaagaa gtgggatcag 720
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taaataaatt tatgaaatga aaatacaaga tgatctacca atctcaciaa tattatagaa 900
aagcttnaaa ttatnggggt cagtgaaaac gctgtgacca tcaactcaca agaacacccc 960
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&lt;210&gt; 404

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-453-330 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-453-330.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-453-330.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 172..189

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 578..597

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-453-330 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 19,31,101,212,337,520,661

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 404

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aatagcaaat tagttcagtg tgttatctag aaaacactgt cactttcaga gcctttcatt 180
gtgcattctca ttttattcct atgaataatt tntgctaaaa ttcattccaat cctaggteat 240
ccaaaaacca gagctttttat aactcatggg ggagccaatg gcatctatga ggcaatctac 300
catgggatcc ctatgggtggg cattccattg ttttttngat caacctgata atattgctca 360
catgaaggcc aaggagcag ctggttagagt ggacttcaac acaatgtcga gtacagacct 420

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351

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taagagagtg attttgaaag aatttaaagtg atttaaccaa tccgaaatct gcttttactt 600
tttatctggt atttaaaaaat tgtatttgaa ccccatatct ctaatgagta accagttagt 660
ngaaacagtt ttctaaataa aaataatttt aaaatgatat agataatata aaaaaatata 720
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ataataaaat gttttaatta aatatctaaa atgtctcaga atataactat tttcttgag 840
aaaaattaat ttttattatt atctttattg taacagactt gaaaatgaga ttttaattttg 900
atagcataaa acccacctat ttatggcaaa aattccaaat atttttacta tgtttacaga 960
gtcatgaagt catcaccagt gtataagttt ggaacatttt t 1001

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&lt;210&gt; 405

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-455-367 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-455-367.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-455-367.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 135..152

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 545..564

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-455-367 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 161,248

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 405

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gaatattatg aaattatcaa gaattcaaca tgatcaacca gtgaagcccc tggatcgagc 240
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cccacaacct cacctgggtc cagtaccact ctttggtatg gattgggttc ctgctggctt 360
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tcttcctgtg acaaaaaaaaa atccttttca agtctacctt gtcaagtaaa aatttgtttt 600
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352

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ggggaaataa aaaataatat aaagccatat gagcttgat tgaaatttgt tgcacttata 720
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ttttttgtag tgatgagatc tcatttgtgt ctccatgctg atttcaaact cctgggctca 900
aacaatcctc ccatttttag atcccaaagg gatgagatta caggtatgta ccaccataac 960
tttacaaaat gagattttta tataagaatg attcaaagt t 1001

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&lt;210&gt; 406

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 12-5-158 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 484..502

&lt;223&gt; 12-5-158.mis1

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 12-5-158.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 346..366

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 801..821

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 12-5-158 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 314,336,793

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 406

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aaccattgag ggccaggaag tggacttcct cctgggcact ggcgcgccct tctcaatttt 180
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gcctgtaacc aggtatttct ctgcctcctc cagctgcaat tgagagactt tgctctttca 300
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tacttgaaga agaaatcaac tttgagctct gggccttaga aggacaattt ggaagggcaa 480
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353

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&lt;210&gt; 407

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-9-367 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..518

&lt;223&gt; 12-9-367.mis1, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-9-367.mis2, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 847..865

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 386..406

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-9-367 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 231,891

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 407

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 aaggctgaat ccaagggact gagaagatac agcctgcagg cccactttc acagcacctc 360  
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&lt;210&gt; 408

354

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<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-10-303 : polymorphic base T or C

<220>
<221> misc_binding
<222> 502..521
<223> 12-10-303.mis1, potential complement

<220>
<221> misc_binding
<222> 482..500
<223> 12-10-303.mis2

<220>
<221> primer_bind
<222> 787..807
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 335..355
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-10-303 potential probe

<220>
<221> misc_feature
<222> 67,85,89,608,657,722..723,752,854,885
<223> n=a, g, c or t

<400> 408
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ctaaacagct gaaggctgtg ttatgggccca ttagtcactc atatttggct cagaataaat      180
ctctttgaat attttacaga gtttgactca tttcctcaca ataatttgac acttgaacat      240
gtgggaactc ccagaaaacc caggaccccc aaaatgtttc ctgaacttga aactaaagta      300
ccagaagggg ccatttgaag gcctttcaaa ctttaagctc ctctgatgga actggtcagt      360
cctcttgagc cctggatctc cctttggttg acagacatca atttttcctg aacttatttt      420
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cttctccatt gccttttata yaaaagataa actcaattgg cttgtctgca catttgcattg      540
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anntaattag tttcttctgt gtttaaatgt tntttaactg ctgagatatt taaaagcttg      780
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ctatggaatt ttnatgact ctccatcaag tataccttca tgttncatat attttcctaa      900
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<210> 409
<211> 1001
<212> DNA
<213> Homo Sapiens

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355

<220>  
 <221> allele  
 <222> 501  
 <223> 12-14-264 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 482..500  
 <223> 12-14-264.mis1

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-14-264.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 237..257  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 680..700  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-14-264 potential probe

<220>  
 <221> misc\_feature  
 <222> 307,577,897  
 <223> n=a, g, c or t

<400> 409  
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 caacattaag catactatat ttatgtaata aatgtatgat tgtgcaccat atgtcctact 660  
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 ataattttaa gaaattttaga aatttctctc atgaaaaaat atacatttcg tgaaaaatat 780  
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<210> 410  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 499  
 <223> 12-17-86 : polymorphic base T or A

<220>  
 <221> misc\_binding  
 <222> 500..518  
 <223> 12-17-86.mis1, complement

<220>  
 <221> misc\_binding  
 <222> 479..498  
 <223> 12-17-86.mis2, potential

<220>  
 <221> primer\_bind  
 <222> 565..584  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 121..140  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 487..511  
 <223> 12-17-86 potential probe

<220>  
 <221> misc\_feature  
 <222> 44,197,388..389,398,407,627,852,856,941  
 <223> n=a, g, c or t

<400> 410  
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 tacgctgact cccctttctg agttagtatt tttcaaaatg tcacatttgt agtttccatt 180  
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 gactgaataa tcatgactwc ttagtctctt ctttgcata ttccccgagt tgtcatttat 540  
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 agggaggaga gagagagaga gagaaangag gtataagtaa aagacataga cagacagaca 660  
 aacggagaca gactgccttt tctaacacag tcttcaaaac tatgctgtca cttccagtat 720  
 actttctttg ctgagaaagc cactgagtctt gccaaaatc aaggggaggg accacgggtt 780  
 gataagaatg tcaaagtgtt tgcagtgatg tctcaaaatt ccacaaatat aacctattga 840  
 gttagtcaac angaangcct ctaagcaact atgaaattca ttacatatgg acattaaaaa 900  
 cacaataaaa tcctaaaaag tgtgaattta tattttttac naactactga aacattaagc 960  
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<210> 411  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-19-163 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 481..500



357

&lt;223&gt; 12-19-163.misl, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-19-163.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 339..357

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 781..801

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-19-163 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 612,814,950

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 411

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tttatttgcg	tacaggtggt	tatagtattc	tctgatggca	gtttgtattt	ctgtgggatc	180
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acatacagca	ttctggagaa	ctttggccca	gccatgaaaa	gtactgttac	ctagctgaca	300
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aatccatatt	caaaataagt	atgtattcca	gtaagaaaaa	aatgcagctc	cttctatgat	600
ggaggtgggc	tnaaggtaat	caatctgcca	ctgggaagct	ggttgattat	cttggagaat	660
gatgccatat	caggggctta	gcatggctct	ctgttcctgc	atattgggca	cacagtgggtg	720
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aattttcttc	tagtgatgtc	attctttgaa	gaacattcac	atgactcacn	aaatgtcttc	960
atatattttg	ccccctcaa	gaggtttatc	cagatagttt	t		1001

&lt;210&gt; 412

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-457-284 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-457-284.misl, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

358

<222> 504..523  
 <223> 10-457-284.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 220..238  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 621..639  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 10-457-284 potential probe

<220>  
 <221> misc\_feature  
 <222> 715  
 <223> n=a, g, c or t

<400> 412  
 gggtcataaaa attattgctt gactagagta attgtaaaca taaaagaaca ccaaacacac 60  
 taaaataaat atgagggtcat caatcttttg ttggtctcct tggcatgcac ctattcagac 120  
 tgttagtatt atgtatttac ttcaaatttt agcagttata ttttaacttg attgattttt 180  
 cctcagatat aagtatgaga aatgacagaa agaaaacaaca actggaaaag aagcattgca 240  
 taagaccagg atgtctctga aatggacgtc agtctttctg ctgatacagc tcagttgtta 300  
 ctttagctct ggaagctgtg gaaaggtgct agtgtggccc acagaatata gccattggat 360  
 aaatatgaag acaatcctgg aagagcttgt tcagaggggt catgaggtga ctgtgttgac 420  
 atcttcggct tctactcttg tcaatgccag taaatcatct gctattaaat tagaagttta 480  
 tcctacatct ttaactaaaa atkatttgga agattctctt ctgaaaattc tcgatagatg 540  
 gatatatggt gtttcaaaaa atacattttg gtcataattt tcacaattac aagaattgtg 600  
 ttgggaatat tatgactaca gtaacaagct ctgtaaagat gcagttttga ataagaaact 660  
 tatgatgaaa ctacaagagt caaagtttga tgcattctg gcagatgccc ttaantccct 720  
 gtggtgagct actggctgaa ctatttaaca taccctttct gtacagtctt cgattctctg 780  
 ttggctacac atttgagaag aatgggtggag gatttctgtt ccctccttcc tatgtacctg 840  
 ttgttatgtc agaattaagt gatcaaatga ttttcatgga gaggataaaa aatatgatac 900  
 atatgcttta ttttgacttt tggtttcaaa tttatgatct gaagaagtgg gaccagtttt 960  
 atagtgaagt tctaggtaag tcatgtgtct aactggtgct t 1001

<210> 413  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 10-460-221 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 10-460-221.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..523  
 <223> 10-460-221.mis2, potential complement

<220>

359

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<221> primer_bind
<222> 283..301
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 686..704
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 10-460-221 potential probe

<220>
<221> misc_feature
<222> 128,707,966
<223> n=a, g, c or t

<400> 413
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gacttatatt aataattatt tatgattgtg aatatactga tgttacatta aagatgtgat    120
ttcttctnac agatctctga atacatttgc cttccttata tatacatatg agcaacatat    180
gcaataaaata aaatctaaat tatgactata tataaatgta tttatatata ttttatcaat    240
gcacagacat tttatatatg tttgggtatg ttattccaag tcctttcagg aaaataacctg    300
catattcaaa taacaattct cgtgttagct accttttgtt ttgttttgtt tttttccatc    360
aggaagaccc actacattat ttgagacaat ggggaaagct gaaatgtggc tcattcgaac    420
ctattgggat tttgaatttc ctgcgccatt cttaccaaat gttgattttg ttggaggact    480
tcactgtaaa ccagccaaac ccytgccctaa ggtaaatgta ttcttgtttc atttgttgc    540
ttgacatttt cagaaggaat ggctggatat gtttctttca gagtgtttaa ctcagagtga    600
ggggaatatg ggaggtcaaa aacaaggact tgccattaga aaatcatata tttctgtagt    660
atcacaagta tgtgaatgtt attatcatta aagaccaaag aggttttact agggagattt    720
tgaaaaacagg gttgggttaaa gtaaggcctt cattgtgcc aatacaaaac acaggtaagt gctggatttt    780
atttcttcaa aaaatatatt tagagtgatt aatacaaaac acaggtaagt gctggatttt    840
cagagaataa aggtagcaca gtttctgctc cctcatgcct tacattgtac tttgaaagat    900
agaataaaaa caagtgaaaa agaaaagtct aaaaagtgtt ataaggaaag accacaatga    960
taagnaaat atgcagaaga gatcccaaac tcattgacaa t                                1001

<210> 414
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 10-460-232 : polymorphic base A or G

<220>
<221> misc_binding
<222> 483..502
<223> 10-460-232.mis1, potential

<220>
<221> misc_binding
<222> 504..523
<223> 10-460-232.mis2, potential complement

<220>
<221> primer_bind
<222> 272..290
<223> upstream amplification primer

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360

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<220>
<221> primer_bind
<222> 675..693
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515
<223> 10-460-232 potential probe

<220>
<221> misc_feature
<222> 117,696,955,992
<223> n=a, g, c or t

<400> 414
aattcaagat gagaggagat ttgggtgggg acagtcaaac catattagtg acttattttta      60
ataattatatt atgattgtga atatactgat gttacattaa agatgtgatt tcttctnaca      120
gatctctgaa tacatttgcc ttccttatat atacatatga gcaacatatg caataaataa      180
aatctaaaatt atgactatat ataaatgtat ttatatatat tttatcaatg cacagacatt      240
ttatatatgt ttgggtatgt tattccaagt cctttcagga aaatacctgc atattcaaat      300
aacaattctc gtgttagcta ccttttggtt tgttttggtt ttttccatca ggaagaccca      360
ctacattatt tgagacaatg gggaaagctg aaatgtggct cattcgaacc tattgggatt      420
ttgaatttcc tcgcccatte ttaccaaagtg ttgattttgt tggaggactt cactgtaaac      480
cagccaaacc cctgcctaag gtraatgtat tcttggttca tttgtttgct tgacattttc      540
agaaggaaatg gctggatatg tttctttcag agtggttaac tcagagtggag gggaatatgg      600
gagggtcaaaa acaaggactt gccattagaa aatcatatat ttctgtagta tcacaagtat      660
gtgaatgtta ttatcattaa agaccaaaga ggtttnacta gggagatttt gaaaacaggg      720
ttggttaaag taaggccttc attgtgccac caaaagata gtatgattca tttcttcaaa      780
aaatatttgt agagtgatta atacaaacca caggtaagtg ctggattttc agagaataaa      840
ggtagcacag tttctgctcc ctcatgcctt acattgtact ttgaaagata gaataaaaac      900
aagtgaaaaa gaaaagtcta aaaagtgtta taaggaaaga ccacaatgat aaagnaaata      960
tgcaagaagag atcccaaact cattgacaat tnaaagtggag t                        1001

<210> 415
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 10-460-235 : polymorphic base C or T

<220>
<221> misc_binding
<222> 483..502
<223> 10-460-235.mis1, potential

<220>
<221> misc_binding
<222> 504..523
<223> 10-460-235.mis2, potential complement

<220>
<221> primer_bind
<222> 269..287
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 672..690
<223> downstream amplification primer, complement

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<220>
<221> misc_binding
<222> 491..515
<223> 10-460-235 potential probe

<220>
<221> misc_feature
<222> 114,693,952,989
<223> n=a, g, c or t

<400> 415
tcaagatgag aggagatttg ggtggggaca gtcaaaccat attagtgact tattttaata      60
attatttatg attgtgaata tactgatgtt acattaaaga tgtgatttct tctnacagat      120
ctctgaatac atttgccttc cttatatata catatgagca acatatgcaa taaataaaat      180
ctaaattatg actatatata aatgtattta tatatatatt atcaatgcac agacatttta      240
tatatgtttg ggtatgttat tccaagtcct ttcaggaaaa tacctgcata ttcaaataac      300
aattctcgtg ttagctacct tttgttttgt tttgtttttt tccatcagga agaccacta      360
cattatttga gacaatgggg aaagctgaaa tgtggctcat tcgaacctat tgggattttg      420
aatttcctcg cccattctta ccaaagtgtg attttggttg aggacttcac tgtaaaccag      480
ccaaaccctc gcctaaggta aaygtattct tgtttcattt gtttgcttga cattttcaga      540
aggaatggct ggatatgttt ctttcagagt gtttaactca gagtgagggg aatatgggag      600
gtcaaaaaca aggacttgcc attagaaaat catatatatt tgtagtatca caagtatgtg      660
aatgttatta tcattaaaga ccaaagaggt ttnactaggg agattttgaa aacagggttg      720
gttaaagtaa ggccttcatt gtgccacca aaagatagta tgattcattt cttcaaaaaa      780
tatttgtaga gtgattaata caaaccacag gtaagtgtcg gattttcaga gaataaagg      840
agcacagttt ctgctccctc atgccttaca ttgtactttg aaagatagaa taaaaacaag      900
tgaaaaagaa aagtctaaaa agtggtataa ggaaagacca caatgataaa gnaaatatgc      960
agaagagatc ccaaactcat tgacaattna aagtgagtac t                          1001

<210> 416
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 503
<223> 10-460-236 : polymorphic base A or G

<220>
<221> misc_binding
<222> 483..502
<223> 10-460-236.mis1, potential

<220>
<221> misc_binding
<222> 504..523
<223> 10-460-236.mis2, potential complement

<220>
<221> primer_bind
<222> 268..286
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 671..689
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 491..515

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362

&lt;223&gt; 10-460-236 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 113,692,951,988

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 416

caagatgaga	ggagatttgg	gtggggacag	tcaaaccata	ttagtgactt	attttaataa	60
ttattttatga	ttgtgaatat	actgatgtta	cattaaagat	gtgatttctt	ctnacagatc	120
tctgaatata	tttgccctcc	ttatatatac	atatgagcaa	catatgcaat	aaataaaaatc	180
taaattatga	ctatatataa	atgtatttat	atatatttta	tcaatgcaca	gacatttttat	240
atatgtttgg	gtatgttatt	ccaagtcctt	tcaggaaaat	acctgcatat	tcaaataaca	300
attctcgtgt	tagctacctt	ttgttttggt	ttgttttttt	ccatcaggaa	gacccactac	360
attatttgag	acaatgggga	aagctgaaat	gtggctcatt	cgaacctatt	gggattttga	420
atttcctcgc	ccattcttac	caaagtgtga	ttttgttgga	ggacttcact	gtaaaccagc	480
caaaccctcg	cctaaggtaa	atrtattcct	gtttcatttg	tttgcttgac	attttcagaa	540
ggaatggctg	gatatgtttc	tttcagagtg	tttaactcag	agtgagggga	atatgggagg	600
tcaaaaacaa	ggacttgcca	ttagaaaatc	atatatttct	gtagtatcac	aagtatgtga	660
atgttattat	cattaaagac	caaagagggt	tnactagggg	gattttgaaa	acagggttgg	720
ttaaagtaag	gccttcattg	tgccacccaa	aagatagtat	gattcatttc	ttcaaaaaat	780
attttagtag	tgattaatac	aaaccacagg	taagtgcctg	attttcagag	aataaaggta	840
gcacagtttc	tgctccctca	tgctttacat	tgtactttga	aagatagaat	aaaaacaagt	900
gaaaaagaaa	agtctaaaaa	gtgttataag	gaaagaccac	aatgataaag	naaatatgca	960
gaagagatcc	caaactcatt	gacaattnaa	agtgagtact	c		1001

&lt;210&gt; 417

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 503

&lt;223&gt; 10-460-285 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 483..502

&lt;223&gt; 10-460-285.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 504..523

&lt;223&gt; 10-460-285.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 219..237

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 622..640

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 10-460-285 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

363

&lt;222&gt; 64,643,902,939

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 417

tattttaata	attatttatg	attgtgaata	tactgatgtt	acattaaaga	tgtgatttct	60
tctnacagat	ctctgaatac	atttgccttc	cttatatata	catatgagca	acatatgcaa	120
taaatataaat	ctaaattatg	actatatata	aatgtattta	tatatatttt	atcaatgcac	180
agacatttta	tatatgtttg	ggtatgttat	tccaagtcct	ttcaggaaaa	tacctgcata	240
ttcaaataac	aattctcgtg	ttagctacct	tttgttttgt	tttgtttttt	tccatcagga	300
agacccacta	cattatttga	gacaatgggg	aaagctgaaa	tgtggctcat	tcgaacctat	360
tgggattttg	aatttcctcg	cccattctta	ccaaatgttg	attttgttgg	aggacttcac	420
tgtaaaccag	ccaaaccctt	gcctaaggta	aatgtattct	tgtttcattt	gtttgcttga	480
cattttcaga	aggaatggct	ggwtatgttt	ctttcagagt	gtttaactca	gagtgagggg	540
aatatgggag	gtcaaaaaca	aggacttgcc	attagaaaat	catatatattc	tgtagtatca	600
caagtatgtg	aatgttatta	tcattaaaga	ccaaagaggt	tnactaggg	agattttgaa	660
aacagggttg	gttaaagtaa	ggccttcatt	gtgccacca	aaagatagta	tgattcattt	720
cttcaaaaaa	tatttgtaga	gtgattaata	caaaccacag	gtaagtgtcg	gattttcaga	780
gaataaaggt	agcacagttt	ctgctccctc	atgccttaca	ttgtactttg	aaagatagaa	840
taaaaaacaag	tgaaaaagaa	aagtctaaaa	agtgttataa	ggaaagacca	caatgataaa	900
gnaaatatgc	agaagagatc	ccaaactcat	tgacaattna	aagtgagtac	tcaataatgt	960
gcagagatag	gtgaaacgat	gaggggttga	taaacacca	a		1001

&lt;210&gt; 418

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-605-58 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-605-58.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..520

&lt;223&gt; 12-605-58.mis2, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 444..464

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 901..921

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-605-58 potential probe

&lt;400&gt; 418

gttcagcaga	acagcagaca	aattactttt	atttttaatt	tcaatactct	ttgcttttaga	60
aacaattttg	ctgtggcact	tcctagtctg	tgactctgtc	ttgaaaaggg	ctggaactca	120
ggctgcgtag	acatacatat	ttcattttatc	tggtcttcaa	cttaaacatt	cacctcttta	180
taaaaacttc	tctgtaattc	tgaaggtaat	tccccttttt	ttgatggagt	tcaaaaattg	240
ctgacttgca	ctgttgggaa	atcatcactg	actgtgattt	cattttccat	actttcatat	300

364

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ttacatccac ctgctttttc tgttcagtg gagacaagca gttggagcca accttctgag 360
cttcttcaaaa aaggctatca acaaaatatt cacaatttgt tgattatcac tgagagattg 420
attgtcagat actgtttcac aaacccatct tcttttatac tttcgagatt atctacctct 480
ttcattctct tttgtcttat kctgggcagg gtctcctggc ctcagcctca ctcagctcag 540
aggccacgag cacccttttg ccccgggctc cgtctggagc ccgctgctcc agacctcctt 600
aggttcgcc gcggcacagc taggatcctg acggactctc taggtctctg ccaggcacca 660
gacaccaaaa gcatgctgat gtagtctcaa ttgatttgtt tttaaattgg taattaaacc 720
tccaccttta ttgaaagatc atttggaaga ttttgttctc atactcggtc agttaataaa 780
catatgattt gtcaaaagat agattttaga aacattttat gaattcaaaa aatttgaggg 840
aaatatattg gttgttttag aaagctctgtc aagatgtagt tttaatcaag aaacttatgt 900
caaaagttta agaataaagc cttaataatta ttcttgacga tgccttggt aaaatctgct 960
ggtggggata attaaaatac acattgtata tactttgtat t 1001

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&lt;210&gt; 419

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-607-207 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-607-207.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-607-207.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 689..707

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 179..199

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-607-207 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 531,558

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 419

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atgtacatgt atatatgtgt tcactttctc caaaacaagg cacagaatta ttgagctgcc 60
ttacagatgt ttgtgaaact gaatgtgttt gtagaagaca gttaaaaatg ataattattc 120
tatataaccc tattttatat aaggatgtct catgtttctc tctttcctca gaagcctgta 180
actctattca tcccataaca ctaatatgaa aacataagtc atttagcatg accaaaaata 240
ggataatgat tttttaaaat gtaagcatat tgcaactaat cataatcagt aacaattaaa 300
agaccacaaa aattaccaaa aattggcaaa taataaaatc tcaggaattc tcaaggccct 360
cttaaaagct acacatcaag gaaacacttt atgatcaaa tagtatgggt taacacaaac 420
ctatgtgtct ttattggtaa gaagaccatc atccaagtct cacaataacc actctatgac 480
ctatggcatg aatgtattct raaaataaat taaaattttg atgctaaggg naattattgg 540

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365

aagagattgc agcaacantt tttataaaat gtatctagat aaatgaaaca tttttaagaa	600
aaattgaatg aatatagtgt catagtagat aatattgcta aatcatgcat acaaacctaa	660
agtaattttac atcaagctta tcctgcaaca catagataaa ggactgggtat ctcttgata	720
gtgagagcta ttatatatca ccaacccaaa agacaaacag accaatatga aaatagatga	780
aggatatgaa ttgacaattc acagaagtaa aaatggccaa tcacatcaaa aagatgggcta	840
cttgtagtag aaagttggac attataattt agtaagtcaa taaccttaag tcaatgaatt	900
tattttaact tttattttaa gctcaggggt acatgtgcaa gtttggtaca taggtaaata	960
tgtgtcaaca cccaggtatt aagcctagta cccactaatt a	1001

&lt;210&gt; 420

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-609-119 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..518

&lt;223&gt; 12-609-119.mis1, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-609-119.mis2, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 597..615

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 114..134

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-609-119 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 454

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 420

ttttaaaaat acttaccat atattggtctg aattgacata ttttagaatt tcttttttta	60
tttgtaaaca gttttataaa gaaattttccc ccaatgatta ggaccagaaa gtatatattgt	120
tatgacagaa ggggttggtc attatttttac tgagaaaaag agaatttaga acaaaaggga	180
caacagcaac aaaaagatga gtatacaatt ttgatattgat atccaggaaa ctatgggtc	240
tacatttctt tctctcttct tctctctctt tttttttttt ttggtgtgag aatacattga	300
gcatgtggc attagagaat ggattttaagt ttaaaaacag ggacacatca ggaaacacaa	360
gtcagaataa ttctcattca tttcaaagca aacacatatt caccaggagc ttcatatagt	420
gtgaggggagc tactctaggg ggtgagcaga tctnccactg gagaaagtcc tgggtgacctc	480
tcccactgtg gttcaagtkc cccctgtgag acacagcaaa gtgatgatga ggggtcccca	540
cattcagtta tacatagcac atcaaattca cagtgtgatt tcaggacaaa aggtgtcata	600
gtcatacctt agcaatgacc tgagaaataa gatcacttaa ttatgtaact atatatata	660
taatactgta ttataaatgg aattctcaga actattttccc agaaattcca aaccacaata	720
ccagactgct gaatgtcagt gattcttata ctccagcttt taagggtgtt ttgggggggtg	780

366

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ggggtgtcgg gggcagtgag gggtagtggg gtaagactag gaaaccctaa ctttaagtga      840
aatataaagg gtttagagagc tagaagctaa ataatgtag atataatatac attccaattg      900
taactaactt ttcctcaatg ctcatagtca tgtaatggct ccaatgactc ctaactaaaa      960
gaactgaaca gaaaaaata aaataaata aaactacccc a                               1001

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&lt;210&gt; 421

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 499

&lt;223&gt; 12-609-180 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 479..498

&lt;223&gt; 12-609-180.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 500..519

&lt;223&gt; 12-609-180.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 658..676

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 175..195

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 487..511

&lt;223&gt; 12-609-180 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 34,53..54,515

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 421

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ggtgacagag aaagactcca tctggaaaaa aaanaaagat gtcataaagt tannttatat      60
tttttaaaaa tacttaccaa tatatggtct gaattgacat attttagaat ttcttttttt      120
atttgtaaag agttttataa agaaatttcc cccaatgatt aggaccagaa agtatatttg      180
ttatgacaga aggggttggt cattatttta ctgagaaaaa gagaatttag aacaaaaggg      240
acaacagcaa caaaaagatg agtatacaat tttgatatga tatccaggaa acttatgggt      300
ctacatttct tttcctcttt ctctctctct tttttttttt tttgggtgtga gaatacattg      360
agcgatgtgg cattagagaa tggatttaag tttaaaaaca gggacacatc aggaacaca      420
agtcagaata attctcattc atttcaaagc aaacacatat tcaccaggag cttcatatag      480
tgtgagggag ctactctakg gggtgagcag atctnccact ggagaaagtc ctggtgacct      540
ctcccactgt ggttcaagtg cccctgtgta gacacagcaa agtgatgatg aggggtcccc      600
acattcagtt atacatagca catcaaattc acagtgtgat ttcaggacaa aaggtgtcat      660
agtcatacct aagcaatgac ctgagaaata agatcactta attatgtaac tatatagtat      720
ataatactgt attataaatg gaattctcag aactatttcc cagaaattcc aaaccacaat      780
accagactgc tgaatgtcag tgattcttat acttcagctt ttaagggtgtt tttggggggg      840
gggggtgtcg ggggcagtga ggggtagtgg ggtaagacta ggaaacccta actttaagtg      900
aaatataaag ggttagagag ctagaagcta aataaatgta gatataatat cattccaatt      960
gtaactaact tttcctcaat gctcatagtc atgtaatggc t                               1001

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367

<210> 422  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 499  
 <223> 12-609-233 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 479..498  
 <223> 12-609-233.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 500..519  
 <223> 12-609-233.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 711..729  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 228..248  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 487..511  
 <223> 12-609-233 potential probe

<220>  
 <221> misc\_feature  
 <222> 87,106..107,568  
 <223> n=a, g, c or t

<400> 422  
 ctgggaggca gaggttgcaa tgagccaaga ttgcatcatt gcactccagc ctgggtgaca 60  
 gagaaagact ccatctggaa aaaaaanaaa gatgtcataa agttanntta tatttttta 120  
 aaatacttac caatatatgg tctgaattga catatttttag aatttctttt tttatttgta 180  
 aacagtttta taaagaaatt tcccccaatg attaggacca gaaagtatat ttgttatgac 240  
 agaaggggtt ggtcattatt ttactgagaa aaagagaatt tagaacaaaa gggacaacag 300  
 caacaaaaag atgagtatac aattttgata tgatatccag gaaacttatg ggtctacatt 360  
 tcttttcttc tttctctctc tctttttttt ttttttggtg tgagaatata ttgagcgatg 420  
 tggcattaga gaatggattt aagtttaaaa acagggacac atcaggaaac acaagtcaga 480  
 ataattctca ttcatttcra agcaaacaca tattcaccag gagcttcata tagtgtgagg 540  
 gagctactct aggggggtgag cagatctncc actggagaaa gtcctgggtga cctctccac 600  
 tgtggttcaa gtgccccctg tgagacacag caaagtgatg atgaggggtcc cccacattca 660  
 gttatataca gcacatcaaa ttcacagtgt gatttcagga caaaagggtg catagtcata 720  
 cctaagcaat gccttgagaa ataagatcac ttaattatgt aactatatag tatataatac 780  
 tgtattataa atggaattct cagaactatt tcccagaaat tccaaaccac aataccagac 840  
 tgctgaatgt cagtgattct tatacttcag cttttaagggt gtttttgggg ggtgggggtg 900  
 tcggggggcag tgaggggtag tggggtaaga ctaggaaacc ctaactttaa gtgaaatata 960  
 aaggggttaga gagctagaag ctaataaat gtagatataa t 1001

<210> 423  
 <211> 1001  
 <212> DNA

368

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<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-611-294 : polymorphic base G or A

<220>
<221> misc_binding
<222> 502..520
<223> 12-611-294.mis1, complement

<220>
<221> misc_binding
<222> 481..500
<223> 12-611-294.mis2, potential

<220>
<221> primer_bind
<222> 776..795
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 251..271
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-611-294 potential probe

<220>
<221> misc_feature
<222> 406,571,773,943..944
<223> n=a, g, c or t

<400> 423
taaatcgtgc acactggacc aaaacttgta agtaccaaat ttcaattaat cttttgagag      60
atgagacaat aaataattca ttcgaatggg ctagatatag ggaaagtaaa tcttgcagag      120
ccattcatct ctaatgtctt ttagcaacat gccgaagcac acaagtaaag ggccactaat      180
ccaaagacaa atgaggaatg tgaaagttaa aaatctcagg accatcaagc aaggtctctg      240
caaagaaaca gtaagtatag tcagcagtca gaaaattggc agacagcaga cagtctttag      300
tgaatgacag agagcacagg gagggctaag tgaaagggag tagtaattaa cattttgtag      360
cattgaaccc atttcttggt gtgaagggct tctaaccctt ttagcntttc ggaaaggcct      420
ctaaacctct tatctgtcag aagggcctct aactgtccta agttgggcct ctaaaccgat      480
tttgaacagt gtctctgttc ragtataaaa atatgttcca ccacttacct aaatcagcca      540
attggtgttg catagtctat ttcctttggg nttggaatct cacctcattt aggtccagc      600
agagtcacca gaggaataa ttactggaaa agtgggtctg tcccagacgt taaggggtggg      660
tttgtggatc tcatgtggga aagaatgcaa ggtgagtcgc agagtacagt aaaattaaca      720
gtttgttaga gactactcta ttacagaata gggcatcctc agaaagcaag aangagaaat      780
acccctacct taaacttagt gcttgcttat atagggtgtt aagaatactg tactttatta      840
caaagggttg tgatcagctt gtgacaggct attaatattg ttattttcct atgtattatt      900
gatttcagca agattttaca agtgggctag ttttttaaag gannaaacta attcttaaat      960
taagaagttt ttgttttcaa actattggga cattttcata a                                1001

<210> 424
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele

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369

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<222> 501
<223> 12-612-41 : polymorphic base C or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-612-41.mis1, potential

<220>
<221> misc_binding
<222> 502..520
<223> 12-612-41.mis2, complement

<220>
<221> primer_bind
<222> 461..481
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 981..1001
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-612-41 potential probe

<220>
<221> misc_feature
<222> 383
<223> n=a, g, c or t

<400> 424
gtagaagctg agaaactagt taggagacat ttcaccagtt gtgacacaat tcttctttaa      60
atatgtcaaa atgggctggg cacagtgggt catgtctgta gggccaacac tttgggagac      120
agaggcagaa tgattagacc acaggagttc cagatgagta tgggcactac agcaagacct      180
catatctaaa aatgtaaaaa ataaaaaata tagccaggca tgggtggtgcg cagctgtagc      240
cccagcttct catgaggctg aggtggagga ttacctgaga ccgaaacgga aaggctgcac      300
tgagccatga tcatgtgact gcactcagcc tgtgtaatgg agcaaaaagcc tgtctcaaaa      360
aaaaataaaa actaaaaata tcnaaaatga tcattcccag attctatttc cactatctta      420
cttatagcac ttagaatggc tcacaatatt ttctgctcta gaaaaacatt aactttccca      480
ccgaaaattc cttttttcat ytttaaaggt atttgtcaat gataaaactc caatttataa      540
accaaacttt ctgtaatgac atacattaaa acattaatat tttatgtcaa ttcaatgaca      600
cttactttga atcacttggt tggcgctttt caaagaccat ccatagactt gatatgctta      660
agcaataaat ttacttttaa tggttgatatc tttatattta tccttcagct ataaagagaa      720
tatcatgaaa ttatcaagaa ttcacatga tcaaccgggtg aagcccctgg atcgagcagt      780
cttctggatt gagtttgtca tgcgccataa aggagccaag caccttcggg tcgcagccca      840
caacctcacc tggatccagt accactcttt ggatgtgata gcattcctgc tggcctgcgt      900
ggcaactatg atatttatga tcacaaaatg ttgcctgttt tgtttccgaa agcttgccaa      960
aacaggaaaag aagaagaaaa gggattagtt atatcaaaag c                               1001

<210> 425
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 499
<223> 12-613-302 : polymorphic base C or G

<220>

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370

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<221> misc_binding
<222> 479..498
<223> 12-613-302.mis1, potential

<220>
<221> misc_binding
<222> 500..519
<223> 12-613-302.mis2, potential complement

<220>
<221> primer_bind
<222> 781..799
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 343..363
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 487..511
<223> 12-613-302 potential probe

<220>
<221> misc_feature
<222> 309
<223> n=a, g, c or t

<400> 425
agaaagtgat agtgccagta gcagggggaa gagagtagca gaataaggac aagggataaa      60
tgactagtag tacaatagtg attattactg atactagcat gatctcggct cactgaaacc      120
tccaccccc aggttcaagc gatctgattc tcctgcctca gcctcccag tagctgggat      180
tacaggcacc tgccaacaca tctgactaat ctttgtgttt ttagtagaga cagagtttca      240
ccatgttggc caggctgggc ttgaactcct gacctcaggt gatccacatg cctcagcctc      300
ccaaagtgnc tgggattaca agtgtgagcc tccgttgttg ttaacttgca gaaggtagac      360
ttgaatccaa gtaaataaatt gtgaaactga ttctctaatt cttttgtaca caaaataaatt      420
gttgcgacaa atttgcttgc ttttccatta tgtattagat tctcagataa tgtttgtata      480
tttcaaaaaga ataagactst tgccaaaaag tatcaagtgt ttgaaaaatg catataggca      540
ttgcctttat aatatactca catgaaactg tacagagaat aaatcatgat ggtaagattc      600
aatatcggtg gcaaatactc attaaaaaaa ctctggaaat aattcaaata tcttacatta      660
agaaatggtt aaaaacttac ataatgtgca taccaagcca ttacaatcct tttttcaaaa      720
caatgtttta ttacactgtc agccgtggta caggtatagt tggaaactaa ggaacaaaat      780
gatgagtaga acaagatcac agtttttttg tagaataaaa aggcataatac aatgagaata      840
aaattttcta aataaacatc acatatgtac attgagttat ataaatagaa ttaaaccat      900
aatgaaatgg cacacatttt aaattgagat agagatatgg gtaaattgatt ttctttttct      960
aaattttcat acttttataa aactacttac attgatcttc t                                1001

<210> 426
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-614-471 : polymorphic base T or A

<220>
<221> misc_binding
<222> 502..520
<223> 12-614-471.mis1, complement

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371

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-614-471.mis2, potential

<220>  
 <221> primer\_bind  
 <222> 952..971  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 424..442  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-614-471 potential probe

<220>  
 <221> misc\_feature  
 <222> 109,479  
 <223> n=a, g, c or t

<400> 426  
 gagctggatt ggtaaagtaa aatgaggaca tgcacgttgc aagagtcaag atagaatgaa 60  
 accaaaattg agataaaatc aaaattgaga taaacacatg aaatatttnc acaaatactt 120  
 cagtgaaca tagcatgaat gattaagacc agtaaaacaa ttagaaataa gagagttttg 180  
 ggaataaagc aggcaatcgg gagagcacta attcagcttt ttttctttt ttaacaaata 240  
 acacttttat attttaatca ctttagtggc aattcctaga gtattagggtg attcctattg 300  
 gggctctatg agtcaagatt cagggagaaa aatagaaaga ttcactagtt ctttctactg 360  
 ctgttattgt ttgttctttt gttcattcat tcagttccat tttatttcat gttatcaatt 420  
 acattttagt gtaaccagtg agtttatttc taatttctag aattgatgtg tttcttttnt 480  
 atgtttatgg ttctcgatc wttccataca tatattatat gaactatttt tcatctctgg 540  
 acatttgaac tatgtttatt ataaaacatc attttgattt attttctata tctataatcc 600  
 tttggtgcaa atgttactat tatacatagt atatagttgt atactattgt aaactattgt 660  
 atattgtata tagttgtata ctattgtata ttgtatatag ttggttattt cttcatttgg 720  
 ttttcattag ttaactttca acaggtattg atttctgcaa acattcttca tattctgcat 780  
 tgtgagaaca tcagaataat ttattgattg cctctgctgt aacctaggca gatcactttc 840  
 attacaattt ggcttttctt gtcatttttt ttacaattta ttgttacaat taatttttat 900  
 agcagaaatt ccctgaactc tgtagataca tttttttcct agttttaagg tctttgtctc 960  
 attttgattc cttccaaaac cacttctttt tatagtcct a 1001

<210> 427  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 12-620-192 : polymorphic base G or T

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 12-620-192.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..522  
 <223> 12-620-192.mis2, complement

<220>  
 <221> primer\_bind  
 <222> 309..326  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 777..797  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 491..515  
 <223> 12-620-192 potential probe  
  
 <220>  
 <221> misc\_feature  
 <222> 461..463  
 <223> n=a, g, c or t

<400> 427  
 tatgagaaaa ctacaagagt caaaatttga tgtccttctg gcagatgccg ttaatccctg 60  
 tgggtgagctg ctggctgagc tacttaacat accctttctg tacagtctcc gcttctctgt 120  
 tggctacaca gttgagaaga atgggtggagg atttctgttc cctccttcct atgtacctgt 180  
 tgttatgtca gaattaagt atcaaatgat tttcatggag aggataaaaa atatgatata 240  
 tatgctttat tttgactttt ggtttcaagc atatgatctg aagaagtggg accagtttta 300  
 tagtgaagtt ctaggtaagt cgtgtgtcca attggtgttt attaagttct aattttcctg 360  
 tgcctttgaa ggtgggctta tataaatata atgtcagaag atagtgtttt tatgggaaat 420  
 tatgaattgc aaatgtaaga tgatctatga gtctcaaaaa nnntatagaa tgttgacctt 480  
 atagaatcag ttagaacctt ggkgccatca ctgctacagg acaccaagag agtcataaac 540  
 cttcaatgta aaacacttat gatttcttta agccatcaca tatcattttg ctatacattt 600  
 tttcatcttt aaaaaagtca atagataact caagaaacat cttcatgaag gcagacatac 660  
 aaattttata tttacacata tttctaaaaa tattatcaat gcaggattga ggaacttgta 720  
 cctgagtacc tcagtttcct catttagaaa ttaaattttg tttttcatat aagaaggatt 780  
 ccttcacagt tgagaaatat agtggctcta ctccagaaac agaagcctaa aacttgagat 840  
 ttctaattgt tataattcc ttcaataaca acttcacaat tatttccttc aaaaactgaa 900  
 atcttggtga aagtgaacat ctaagtttta atctatattt tattaactg catctctcca 960  
 tcaaagaaaa taggggccaa aataaggaag agcacatatc t 1001

<210> 428  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 503  
 <223> 12-621-49 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 483..502  
 <223> 12-621-49.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 504..523  
 <223> 12-621-49.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 455..473



373

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 907..927

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 491..515

&lt;223&gt; 12-621-49 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 634

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 428

aacttgga	aa	tggtttgtga	gatataaaat	caaaaataaa	agcaaccaaa	ttaaaattga	60
ttagttggac	ttcatcaaaa	tcaatacttt	tgtgcttcaa	tgaacaccgt	taagaaagta		120
aaaggacagc	ccaaagaata	aaaaaaaaac	aaggaaatca	tatatgtgat	aaggttctta		180
tatcagaaag	atattaaaaa	cacaagtcca	taagaagaca	aataattcaa	ccaaaattag		240
gcaaaacatg	ctcaacatga	tttcatatca	aagaaaggta	aagcataacc	agatatcact		300
tcattcacac	tggctataat	caaacgaaat	ataatcccaa	gtgttggcaa	ggttgggaag		360
aaattaaaa	tcttatacat	tgctgggtgag	aatggaaatt	gaagcatcca	ctgtgaaaaa		420
caatctggaa	attcctatgg	taatttagaa	atttatgggt	aacagacaat	tcctatggta		480
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atagatggat	atacgatctt	cagaaagatg	cattttgggc	atatttctca	caagcacaag		960
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&lt;210&gt; 429

&lt;211&gt; 710

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 364

&lt;223&gt; 12-622-325 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 344..363

&lt;223&gt; 12-622-325.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 365..384

&lt;223&gt; 12-622-325.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 40..59

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

374

<222> 551..569  
<223> downstream amplification primer, complement

<220>  
<221> misc\_binding  
<222> 352..376  
<223> 12-622-325 potential probe

<220>  
<221> misc\_feature  
<222> 333,685,688  
<223> n=a, g, c or t

<400> 429  
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gcactaacct ggtatgacct ctacttatcc tgtcttccac cctcaccttc tacttatcct 180  
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acctttgggt cctcatccaa tctaccctg atatttgggtg caagattcag aagcttgaca 600  
atggccctga aacccacaa tgacaccttc ttaatttagt cttcaaagtc tttaacagtt 660  
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<210> 430  
<211> 1001  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 501  
<223> 12-624-82 : polymorphic base T or C

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-624-82.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-624-82.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 562..582  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 114..134  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-624-82 potential probe

<220>

375

<221> misc\_feature  
 <222> 464..639  
 <223> n=a, g, c or t

<400> 430  
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 gtcatataacc cttcctgtga aatttagcaa attttgtggc tgtctcaaaa aagagaatat 180  
 agtggaagtg aggagcttaa cttccaaagc tcaaaaaaat tgtaataca ttttctctgt 240  
 ctgtccttct gtccttttct ctgtgtttct gtcattcttt tttccttttt ctctgtctct 300  
 ccactcgctt tggagccctg tgctatatcc acagtcacca ggagaaatat accatttggg 360  
 gtagaaatac atatgtatgc ctaataagtc acaattggcc cagcactgca gtagcttgag 420  
 acttttaata ctaatcacca gatatgtgag tgattaacct tcangatgat tagccaagc 480  
 taccttagga cagtaaccaa ytgagaaatg ctgtgccata aacacctaatt tgagcctggc 540  
 caacaacctt gaattaaaac agatgatgat aaaatgttgc tgttctaagc accttcattt 600  
 cagagcagtt ggttaagcag tgatagacaa ccagaaaana tatttttaca tatttgagtt 660  
 atactcctgt aataatatta aaattaatat atacctacaa aaaaagtga tctatattac 720  
 attttgtaac tacctgtaac ccagaaagag tagtcagcat aggatggtgg gtaagagctt 780  
 gcactctgaa tccagataac ctatatcatc ttgcaaaac cattacagaa tcatgtatag 840  
 ttaagcaaat tacttatcag gtctatgcct cacattctcc ttcagtgaat tgtagatgaa 900  
 gcaatgtcac tactactta atagcattgc tgagactaaa agagttgctg tatgtaattg 960  
 ctgtatatag caaaattcaa tttgctataa ctattattat t 1001

<210> 431  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-624-83 : polymorphic base G or A

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-624-83.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-624-83.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 563..583  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 115..135  
 <223> downstream amplification primer

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-624-83 potential probe

<220>  
 <221> misc\_feature  
 <222> 465..640  
 <223> n=a, g, c or t

376

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<400> 431
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agtcataata ccttcctgtg aaatttagca aattttgtgg ctgtctcaaa aaagagaata      180
tagtggaagt gaggagctta acttccaaag ctcaaaaaaa ttgttaatac attttctctg      240
tctgtccttc tgccttttcc tctgtgtttc tgtcattctt ttttcctttt tctctgtctc      300
tccactcgct ttggagccct gtgctatatc cacagtcccc aggagaaata taccatttgg      360
tgtagaaata cataatgtatg cctaataagt cacaattggc ccagcactgc agtagcttga      420
gacttttaat actaatcacc agatatgtga gtgattaacc ttcangatga ttagcccaag      480
ctaccttagg acagtaacca rctgagaaat gctgtgccat aaacacctaa ttgagcctgg      540
ccaacaacct agaattaaaa cagatgatga taaaatgttg ctgttctaag caccttcatt      600
tcagagcagt tgggttaagca gtgatagaca accagaaaaa atatttttac atatttgagt      660
tatactcctg taataatatt aaaattaata tatacctaca aaaaaagtga atctatatta      720
cattttgtaa ctacctgtaa cccagaaaga gtagtcagca taggatgggt ggtaagagct      780
tgcactctga atccagataa cctatatcat ctttgcaaaa ccattacaga atcatgtata      840
gttaagcaaa ttacttatca ggtctatgcc tcacattctc cttcagtga atgtagatga      900
agcaatgtca ctacctactt aatagcattg ctgagactaa aagagttgct gtatgttaatt      960
gctgtatata gcaaaattca atttgctata actattatta t                               1001

```

&lt;210&gt; 432

&lt;211&gt; 989

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 489

&lt;223&gt; 12-624-107 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 469..488

&lt;223&gt; 12-624-107.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 490..509

&lt;223&gt; 12-624-107.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 575..595

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 127..147

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 477..501

&lt;223&gt; 12-624-107 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 477,652

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 432

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tttaaaattt taatgtctta tgtcaacatt ttatgaattc tacagtagtt tagtgatttc      60
ttctaatttt taaggttata tttgcagtaa ttttgagatg tgattacaaa tgcattggca      120
atcctctcat caagtcatat acccttctctg tgaaatttag caaattttgt ggctgtctca      180

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377

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aaaaagagaa tatagtggaa gtgaggagct taacttccaa agctcaaaaa aattgttaat   240
acattttctc tgtctgtcct tctgtccttt tctctgtgtt tctgtcattc ttttttcctt   300
tttctctgtc tctccactcg ctttgagacc ctgtgtctata tccacagtcc ccaggagaaa   360
tataccattt ggtgtagaaa tacatatgta tgcctaataa gtcacaattg gcccagcact   420
gcagtagcct gagactttta atactaatca ccagatatgt gagtgattaa ccttcangat   480
gattagccya agctacctta ggacagtaac caactgagaa atgctgtgcc ataaacacct   540
aattgagcct ggccaacaac ctagaattaa aacagatgat gataaaatgt tgctgttcta   600
agcaccttca tttcagagca gttgggtaag cagtgataga caaccagaaa anatattttt   660
acataattga gttatactcc tgtaataata ttaaaattaa tatataccta caaaaaaagt   720
gaatctatat tacattttgt aactacctgt aaccagaaa gagtagtcag cataggatgg   780
tggttaagag cttgcactct gaatccagat aacctatc atctttgcaa aaccattaca   840
gaatcatgta tagttaagca aattacttat caggtctatg cctcacattc tccttcagt   900
aatgtagat gaagcaatgt cactacctac ttaatagcat tgctgagact aaaagagttg   960
ctgtatgtaa ttgctgtata tagcaaaat   989

<210> 433
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-624-146 : polymorphic base T or C

<220>
<221> misc_binding
<222> 481..500
<223> 12-624-146.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-624-146.mis2, potential complement

<220>
<221> primer_bind
<222> 627..647
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 179..199
<223> downstream amplification primer

<220>
<221> misc_binding
<222> 489..513
<223> 12-624-146 potential probe

<220>
<221> misc_feature
<222> 529,704
<223> n=a, g, c or t

<400> 433
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tttaagggtta tatttgcagt aattttgaga tgtgattaca aatgcattgg caatcctctc   180
atcaagtcata atacccttcc tgtgaaattt agcaaatatt gtggctgtct caaaaaagag   240
aatatagtgg aagtgaggag cttaacttcc aaagctcaaa aaaattgtta atacattttc   300
tctgtctgtc cttctgtcct tttctctgtg tttctgtcat tcttttttcc tttttctctg   360
tctctccact cgctttggag ccctgtgcta tatccacagt cccaggaga aatataccat   420

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378

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ttggtgtaga aatacatatg tatgcctaata agtcacaaat tggcccagca ctgcagtagc 480
ttgagacttt taatactaata yaccagatat gtgagtgatt aaccttcang atgattagcc 540
caagctacct taggacagta accaactgag aaatgctgtg ccataaacac ctaattgagc 600
ctggccaaca acctagaatt aaaacagatg atgataaaat gttgctgttc taagcacctt 660
catttcagag cagttgggta agcagtgata gacaaccaga aaanatattt ttacatatatt 720
gagttatact cctgtaataa tattaataatt aatatatacc tacaaaaaaa gtgaatctat 780
attacatttt gtaactacct gtaaccagaga aagagtagtc agcataggat ggtgggtaag 840
agcttgcaact ctgaatccag ataacctata tcctctttgc aaaaccatta cagaatcatg 900
tatagttaag caaatactt atcaggtcta tgccctcacat tctccttcag tgaaatgtag 960
atgaagcaat gtcactacct acttaatagc attgctgaga c 1001

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&lt;210&gt; 434

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-624-288 : polymorphic base T or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-624-288.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-624-288.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 769..789

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 321..341

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-624-288 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 113,671,846

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 434

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gttttaatat aatattatca atccattagt gccatttaga aaacttaaata tgaccagcaa 60
tccggtgctc tctatctgat aaatttacag cctgtttttg gacacctact tgnatatttt 120
aaatatttat taataaatat attagaatgt taattaaaaa ttttaattat ggctaaaaatt 180
ttagtaaaca aatttttaaa attttaatgt cttatgtcaa cattttatga attctacagt 240
agttagtgta tttcttctaa tttttaaggt tatatttgca gtaattttga gatgtgatta 300
caaatgcatt ggcaatcctc tcatcaagtc atataccctt cctgtgaaat ttagcaaaatt 360
ttgtggctgt ctcaaaaaag agaatatagt ggaagtgagg agcttaactt ccaaagctca 420
aaaaaattgt taatacatatt tctctgtctg tcttctgtc cttttctctg tgtttctgtc 480
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gtccccagga gaaatatatac atttggtgta gaaatacata tgtatgccta ataagtcaca 600
attggcccag cactgcagta gcttgagact ttttaacta atcaccagat atgtgagtga 660

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379

```

ttaaccttca ngatgattag cccaagctac cttaggacag taaccaactg agaaatgctg   720
tgccataaac acctaattga gcctggccaa caacctagaa ttaaaacaga tgatgataaa   780
atgttgctgt tctaagcacc ttcatttcag agcagttggt taagcagtga tagacaacca   840
gaaaaanatat ttttacatat ttgagttata ctctgtgaat aatattaaaa ttaatatata   900
cctacaaaaa aagtgaatct atattacatt ttgtaactac ctgtaaccca gaaagagtag   960
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&lt;210&gt; 435

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-624-293 : polymorphic base T or C

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-624-293.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-624-293.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 774..794

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 326..346

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-624-293 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 118,676,851

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 435

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cttgggtttt aatataatat tatcaatcca ttagtgccat ttagaaaact taaattgacc   60
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attttaataa tttattaata aatatattag aatgttaatt aaaattttta attatggcta   180
aaatttttagt aaacaaattt ttaaaatttt aatgtcttat gtcaacattt tatgaattct   240
acagtagttt agtgatttct tctaattttt aaggttatat ttgcagtaat tttgagatgt   300
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ccacagtcct caggagaaat ataccatttg gtgtagaaat acatatgtat gcctaataag   600
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agtgattaac cttcangatg attagcccaa gctaccttag gacagtaacc aactgagaaa   720
tgctgtgcca taaacaccta attgagcctg gccacaacc tagaattaaa acagatgatg   780
ataaaatgtt gctgttctaa gcaccttcat ttcagagcag ttgggttaagc agtgatagac   840
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380

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 agtagtcagc ataggatggt gggtaagagc ttgcactctg a 1001

<210> 436  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-421-135 : insertion T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-421-135.mis1, potential

<220>  
 <221> primer\_bind  
 <222> 367..385  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 797..817  
 <223> downstream amplification primer, complement

<400> 436  
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 agaacttgcc ttaaaataat ctgggcctaa tgtttttctg gtgggaaatt tttaaattat 180  
 gggatttgca ttctttaatg attattggac taggcaggct ttttgttaca tcttgaatca 240  
 attttggtag gccttggtcc ttgagatgat aacattggtc tcttgattat ccacttgctc 300  
 caaggagggt ttacagact actccctagg cttgaaagcg tgggggtgaat accaccggga 360  
 gtacacgaga taattgtaag aggtgtgtag acttggtttt aaataacatt gaactacatg 420  
 atgagaagtc attctcattt caattatctt tgaatcctta tgggttaagtc aagtggaaaag 480  
 tattgggttg gagccagcat gtccacaccc tcaaaattaa tgtttatatt tatcaaagta 540  
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 aaattttaga cattgtcagt tggcttcttt ttatggaaga aataatttca tgctccttta 780  
 cttcttcttt tccttacctc atccttccag taacattata tcaactctgtt aggttaaatac 840  
 aatagccaat gtttataatt attgtgatta agaaaataat gttgaaagca gggccttata 900  
 gtatactgtg gttacatttc cttcttgtgc aatttttttt ccctaggggt aataattgct 960  
 ctttatttct ttgtttgctt agtttcatgc ccaaactctt t 1001

<210> 437  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-442-133 : insertion C

<220>  
 <221> misc\_binding  
 <222> 502..520  
 <223> 12-442-133.mis1, complement

<220>



381

<221> primer\_bind  
 <222> 616..633  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 184..203  
 <223> downstream amplification primer

<220>  
 <221> misc\_feature  
 <222> 94,734  
 <223> n=a, g, c or t

<400> 437  
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 cccattcat ttaactatct gcacacagag acangaagcc agaaatctga ctgggaagaa 120  
 attcctaccc ttttgccagc atgctaagct tctgggttct ctttccctga gtggccctag 180  
 tgatctggct tgtggcacia ctgcctttgg gggccaagcc gcatcataaa ggaaaagtat 240  
 tcttttttgt tctggccaaa gcaaaatacg cgtaataaaa catagatatt aaccaggctg 300  
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 tttaatctaa cctccgacta ggagtttcaa catgtgtgtc ctgggcaaga tggtcgccct 420  
 gagtaataga aaagaaagag aaaggagaga gagaaaaaca ttgcctgtgg cagggcgggg 480  
 aagggtgaaat gatcagggag gcagagaaag aaccacccat tgcagcgaca ctaaaaagtt 540  
 caggtggctg ctgtcgggtg agcaaggatc ttttccagta atcctaccag ctctcaaatt 600  
 tcccttggtt gggaggaaaa agctcccat gtcccaggat cctgtacatt cctaattctg 660  
 tccccatag ccacagcaa agtacaagg agattaatcc aaagagaata gcagttaaca 720  
 tcccatagt ccgnaacctg ttcttagccg agagggactt taccgaagag gggcctctaa 780  
 cccgctaaat cttagaagg actctaactc tcctaagtcg ggctctaac cagaggtcag 840  
 tcaagcatcc ttgcctttta ttaaggggga gcctttaacc acctctatct taggagagac 900  
 tctaactcca ctaagctggg cctctaacc aatcccatc tttaccagg taccacacca 960  
 cttacccaaa gtcttccaat cagtgtgca gtctatttc t 1001

<210> 438  
 <211> 756  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-449-63 : insertion AT

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-449-63.mis1, potential

<220>  
 <221> primer\_bind  
 <222> 546..563  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 86..106  
 <223> downstream amplification primer

<220>  
 <221> misc\_feature  
 <222> 123,277  
 <223> n=a, g, c or t

382

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<400> 438
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agaaataaat actcataatt attacaattt tcctctgaaa gtagtcacat ggttatacct    120
ggntatattt ttatttttta ttattattat tattatttga gacagggtct caccctgtca    180
cacaggctgg agtgcagtgg tgcaattatg gctcactgct acctctgcca cctgggctca    240
agccatcctc tcaccttagc cttctgaaca gctgggnata catgtgcact aatttttttt    300
tttttgaaac aaaaatattt gtgtagaagg cacaaaagct acaatcacag actccactgt    360
gcaaaggcgc aacctgcctc attgatctct agtgtacacc aaccagctt ccctttccat    420
tcagcctgtg aaaggagata gtgcttgggc catttggtta aagaagggga tgggagatga    480
tcaaaacccc aagtaagggt catccaatat ggtgtctaag cagcaaatga ctaattgctg    540
aagaaggaga ctagacagag gattagaggc agccatgggg ccagtgcagc tgtggagagc    600
tctgagcaaa gaaacaaggt tggcagggtga ggaggcctag gatagaggcc agaaggccaa    660
gcctggggct gcatgtgcac taatttttgt atttttttgc ttttttgata gagatgagat    720
ttcatcatgt tgcacagggt ggtcttgaag tcctgg                                756

<210> 439
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-454-242 : deletion AT

<220>
<221> misc_binding
<222> 481..500
<223> 12-454-242.mis1, potential

<220>
<221> primer_bind
<222> 260..279
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 755..773
<223> downstream amplification primer, complement

<220>
<221> misc_feature
<222> 795..800
<223> n=a, g, c or t

<400> 439
caaatggaaa aaatgaaaac tgtcacactt cttacataac ttcttttatt atctgtattg    60
taaatacaga gtctcaatta ttgactaatt tatactaata gtgggaagca gtgtctagat    120
aatcccagat gattactaat ttaaaactac tttacatttt atttagtaat gtgtttggaa    180
cagttacatt ttgtagctac ttacaatttt actttaactg aaataataag acaaggccat    240
aaatattggc ctccctggct ggggagttgt gtgggtttcc tttccaggca gttgaggttt    300
tccatggagc ctactctaca gtgtcagcag gagtataaca tagaaggctt ccaggtcagg    360
tggcccagag tcaaatcaaa gccccatccc tttttagcaa attttcagc cttacttagc    420
tgtgccggcc tgccctcttc taaaatgagg aaaataataa taccacttc actggtttat    480
tgagaggatt aaatgaggac atgtgttatt tcacaccatt attgcttctg ttgttattat    540
tttaaaatct aggttggtga ttgcatcagt ttcttagggc tgctctaag aaagtaccac    600
aagctgagtg acttacatag cagaaaagtg ttttcttaca ggtaagagg ctggaagtct    660
gaaatcaagg tgtcagcagg gccatgctcc cactgaaacc catagcgggg aatcctttct    720
tgcttctaag ttctagtgt ttcttggct tgtagatgtg tcacttcagt cacacggcca    780
tctttctctt tcttnnnnnn cttcttcttc ttcttctctt tcttctctt cttcttcttc    840
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cttcttcttc ttcttctctt tcttctcttc cttcttcttc ttcttctctt tcttctcttc    960
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<210> 440  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-463-230 : deletion CAT

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-463-230.misl, potential

<220>  
 <221> primer\_bind  
 <222> 255..272  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 773..790  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_feature  
 <222> 136,400,713,730  
 <223> n=a, g, c or t

<400> 440  
 aatcaatgca taaacaaaaa ggatgcccga gaaaacctgt gcttttcaga cacacacaca 60  
 catacacaca cactcactca ctactcact ctgaattggg tcaagtaaaa aaaaattttt 120  
 tttaaaggaa ggaatncttt agcccaactg ggatttcgac gaaattgtat atatcaaaag 180  
 atagctacat ccaacctctc accagaatcc agaggcttct tcctgcaaag atactattga 240  
 aagttttatt agtaagtttg agaaggcgaa agtatgtatc agagggtgaaa gctaacatga 300  
 caattgcttg gccaaagagc gcagcagttt tatgttattg agcagcacag gatgaatcga 360  
 acaccacga tggctagaca gaggcgcctg gctgtgactn ccagagggtc aggcaagcca 420  
 tctgacctct ctgagtcctc acccctgaa atgaaaacca tactcataag actccttgac 480  
 aatggaaaatt ttggagggtga catagttaat gaagtgtcat agaaaggaaa tagctaagaa 540  
 tactctaaaa cctccccatt cccagtatgg gcaatttggg aaaaatgagt aactaaaaat 600  
 taggggttct caaagtgttg ttccaggatc aaatgcatta gtccccacct gggagcttgt 660  
 tagaaatgca aattcccagg ctccctcta ggctcctga attagaactt ttnggggggtg 720  
 tttaacaan gctctccagg tggttacgat gtcccataga gaatcattgg ggcaataat 780  
 cctgtgccct gattagattg caaagctgca ctttcaaaag caagtcaggg tgctgagttg 840  
 cttagtttct attgcaaaat ttctgcttt ggaacaaaat tttcaaaatt ttcagaggggt 900  
 tgctcaggcc agtgatgttt gcaatgcaaa cccaagaatt ctgaaaggag cccatggctg 960  
 aaaggagcca attcttccag gacggactac tccagttgca a 1001

<210> 441  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-462-199 : deletion ATGTCCCACAGCTG

<220>  
 <221> misc\_binding  
 <222> 481..500

384

&lt;223&gt; 12-462-199.mis1, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 303..322

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 736..756

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 255

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 441

caaaagcaag	tcagggtgct	gagttgctta	gtttctattg	caaaatttcc	tgctttggaa	60
caaaattttc	aaaattttca	gaggggttgc	caggccagtg	atgtttgcaa	tgcaaaccga	120
agaattctga	aaggagccca	tggtgaaag	gagccaattc	ttccaggacg	gactactcca	180
gttgcaactg	tgctgtggag	gctgaggagc	atccatgact	gaggaagggc	gagtccccta	240
taatgccctt	gctangggga	gcatcagaca	tccctagact	tacaagtccc	tcgtgtaata	300
aggtgaacta	tggaagccta	tgatttgagg	aaagttccct	ctgattgaaa	gagttcctca	360
aatatgaaga	atgtaacact	tatggggcac	ctactcattg	caaaacactc	tactaggggc	420
tttcacctgt	atgtttaact	cattttgttg	atTTTTTTTT	ttttaagccc	aaatttctga	480
gaggtttaagg	aacttgtctg	gtggtggcgt	ctctgggatt	ctgacttttc	cactgaccat	540
aacatgcac	tggtctaaat	tgagcggacc	acccaggagg	acgttggaat	aaaaagaaat	600
gggaaggggg	tggggagagg	aatgggcata	gaataaaaag	gtagaaaccc	ttgggatcaa	660
aggatgcaac	tgaaagagca	ccaaaatatg	catcattttt	tttaatatgt	tgtctggtac	720
ccaaaattgg	gatgcctctg	tggtatcaac	tttgacattt	aaagaaaggg	cataagatta	780
gatcattcaa	ccataagtgg	aaggaagaaa	caaaataaaa	taaaacctta	caattccctt	840
tgaagtgtgat	caggtgtcaa	ggcagtgatt	tggttttctt	ttacaaaactt	tctttttcca	900
gaaaaatttt	tactgtgggt	ttggagaggt	gtttacaata	aattcatagc	agagctccaa	960
aagggccccac	cccatcta	atcctctgtc	tcagcagatt	a		1001

&lt;210&gt; 442

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-430-287 : insertion T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-430-287.mis1, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 215..233

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 617..635

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

385

&lt;222&gt; 938,965,972

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 442

cgaacaatat	aattatthaa	attatatthtt	gcatggccat	ttgttaatgc	tagttctcac	60
atgtctgtaa	aattaggatt	tttaatatca	attaaatgaa	ggactttgtt	ttgttccactg	120
ttacatctcc	attgcctaga	acattgacat	ataaatattht	gttaaaccgaa	tgaatgaaat	180
aacattctga	caattgaaga	agagattthta	agaatagctg	taacacatca	tacataaaaa	240
tgtaagcat	aaatatttgt	aatatatgta	ccgaaattthta	ttgaccctca	catgtttggc	300
agattcttaa	ttttttctta	ctggctttac	ttagggacgt	gcttccttaa	cacaagagaa	360
agagaagtht	gcccatgaac	atgaagaaat	gaagaaccta	gaagccattg	ttcaagaaat	420
aaaaccaact	gccctcatag	gtaagcattt	tctcctctcc	cttctccccct	accctttatt	480
ttgacctgat	tgthtttttt	actaccaatt	tttgagggtat	aatttagcct	ttttgaattc	540
aaaaaactac	acgttcgctc	aagcgacagt	accctgtaga	gacctagatt	tttaactthta	600
gtcctgtcgt	ctttctctta	ctgatctgag	atgaagcgta	tggccccctc	ctctaactca	660
gcctgttaat	acctggcacg	ttcagcactt	ttctaggctt	gattactcca	gatagacagt	720
ggttctcaaa	taagagggt	ttttgccctg	caggaaacat	ttggcaatgt	ctggaagctt	780
tgagttgtct	taactagggt	agcgggggtg	gaagggagta	aagggagggg	ctactgacat	840
ctagcagata	gaggcagcgg	aatgattcta	ctaaatatcc	tgtagtgcac	tggacagccc	900
ctgcgacaag	gaattatcca	gcctaaaatg	tcaatagntg	ctgaggttga	gaaatactgc	960
agtangataa	gnaccctcct	ggaagacagg	gatcacatct	c		1001

&lt;210&gt; 443

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-718-432 : deletion T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-718-432.mis1, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 913..932

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 479..499

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 346,853..854,883..884

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 443

gtaaccttgg	aaaaattact	taccttcctc	aaggcttaat	tttcttgctt	gtaaattggg	60
gtaaatgct	aatagtacct	tcagtagtac	cttcagacag	ttgttatgat	gattgagaga	120
aaacatataa	aacaaattac	ttttatcaag	tgctataaat	agtaactata	atthtttattt	180
ttcaggaagg	attactagga	ccataattthta	gttaaaagat	gtaggctggc	aaatttggtt	240
taatattaaa	tagagcattt	aatagttgga	attgtcccca	aagaaaataa	tgtatataaa	300
agtacttgca	cataaaaaatt	actcagaggt	tgttattcct	tattgnttht	tgthgttaat	360
attattttgt	aactctccat	cactctgaga	acatggacag	atgtgcttaa	agatataaag	420
atthctactt	ttatgtgaga	tgatggatta	gaattaaatg	tttctaaagt	gttccaactt	480
taggattcta	tgaaagtgg	aaaaaaaaaa	tcgtatttht	tttcaacatt	caggactgg	540
tgthttcataa	aattcagtg	ttcagccatg	cttgtgatca	ttatgagtca	gttgaaattg	600

386

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atcatcacaa tatatttgac ccagaggaat aaacttctca ttcaacattg taaattatTT 660
tgTTTTgcat aacaaaacaa ctttaataaa cttcagtaag ctttagcttt cccttgctac 720
ccaaagtTTa ttTcataaaag agctcaagag acaattggca agttaaataa ttcaagagaa 780
aaaatgTTca caagaagTTt ttatttgctt tgcattctat tccctatgct acatattaat 840
tgtacagtta gcnncaaagt aagTTTTgta gtaaaaatta ctnncaaat aaaaaaggta 900
catagataca agcaactagc aaattaactt cgcaaagaat aagatgtggg ttTacataag 960
ccatttatgc cagttaacta gaccagtTct caactctctc t 1001

```

&lt;210&gt; 444

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-269-301 : deletion T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 482..500

&lt;223&gt; 12-269-301.mis1

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 204..222

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 634..654

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 626

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 444

```

acacagaggc accatctgtg tcaactctgtg tgtggctctgc catgttggtgg ggtgggcact 60
acagactcag gcagctgggc agacaatacc ttagccttag atgatgctga tgcagcccag 120
gagtcagaaa ctgtagtgca gacaatgccc tccttagggc aacacaatta agtgcaatag 180
atgactggct tttctgttag cctcttcatt ggaacccaaa gcagcattac tctaccaaac 240
agagggggagc tggaaagaaa ctacacagtt tgcccagcct agcctctgcc ttgacacgga 300
accatgtgag tctagacatt cacctagatc attccttggg gaccaatgct gctgacacat 360
taactcaata gtttgcctcg gcctgagagg tcatgtaact tgtagaaagt ttagaagcag 420
agattagtgt catttatTTg ccatggctgt gacaacaaaag gaaggaaacag gagtgggaaa 480
acccaaggcc accctggTTt tggtagatgg tgcacacgct tccactaact gttctggggc 540
aaagatccaa atgcactatt gggcctggct atgctgcttc tgctgggtcc cccaaacatg 600
agcctccacg ccatttctca gttgtnattt taccacatat tatcacagtc accggatttg 660
tacagaatat ttggaaccta tactgtctta agggctaccc tttaaagaag agaaaacaag 720
gttttaattc aactgtctgg aacattttat gtttacttat gtggaatact acatcttttg 780
ttataaacag gagggaatgt ggacattcga agggccctac cttttagcta aaagcccata 840
tgaagcatat ggatccattt atacacacca tgcttttcag ctacattttc ctaatttgcc 900
tctctggggc caaccttTgt ggactagcag attcatgggt gagttgaagg atggtgacct 960
cttccatcag cttttcttct tctccagtc ttccaaccct c 1001

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&lt;210&gt; 445

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

387

<221> allele  
 <222> 501  
 <223> 2-13-398 : insertion G

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 2-13-398.misl, potential

<220>  
 <221> primer\_bind  
 <222> 104..121  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 506..525  
 <223> downstream amplification primer, complement

<400> 445  
 tgagctccca tcccgcatcc tgtgtggcac tctgtccatc aagccagtg tgaaggagtt 60  
 cacggaaacc tcagctgtgt ttgaggatgg gaccatgttt gaggtatcg actctgtcat 120  
 ctttgcaaca ggctatgatt attcctaccc cttccttgat gagaccatca tgaaaagcag 180  
 aaacaatgag gttaccttgt ttaaaggcat cttcccccca ctaatggaga agccaacctt 240  
 ggctgtgatt ggcttggttc agtcccttgg agctgccatc cccacagcag acctgcaagc 300  
 ctggtgggct gctaaagtat ttgcaagtag gtggggccatt ctgtctttca ttcattttat 360  
 caatgaacat ttactgaaca cctgctatat gcaaagcact gtgctaggga tacaatgaga 420  
 acaagacaaa catgttcctt gacctctcaa ggcttaaaat ggggtgtggg ggatgccata 480  
 ataggggaaa tttggggggg ttctagttag gggagttgga ctgttgcaaa gagcaaacag 540  
 tatacaggaa gtcataaagg tgagggaag catgaaatgt gtaaggaccc agaaacattt 600  
 tgggtggaag gaataaaag cagaggcagg gagtggcaag aaatataggt ttataagcca 660  
 cgtaaagag cttaaacttc tcatagggat taaggacttc gcaagatttt aagcaagaaa 720  
 aaaatagcag aggataactg caatgtcagg ctacattata aagattggaa gggccctggt 780  
 gagggttgga ggtgtgccag aaacctcact ggtgtcaact tctgtcagaa taacaaagtc 840  
 aggccactct gattctcatg acaatcttct tcttctctcc ctctactcta gacctcatgg 900  
 tctccagggg ctacaagtat gcttatgtga ggaaatcaag aatatgagga ttacatggag 960  
 aaaggcaatg tctcaaatat attaatctac tccagtcata c 1001

<210> 446  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 2-28-132 : insertion T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 2-28-132.misl, potential

<220>  
 <221> primer\_bind  
 <222> 370..387  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 769..793  
 <223> downstream amplification primer, complement

388

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<400> 446
tattcagggg gttaaaggaa gacaaagggt tttaaatggg gaaaaaatac aattacataa    60
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agttactgag cagatgttct tgtagaagtc atttttgtgt aagattatga tggctcttgt    180
gtaagggtgt gggttttgtg gtttttgtta tcaggcacac atcatgagaa cccgctcttt    240
ctggcctttc ccaattctat ttgtcgggtt tcttaacatt agtgactcca tctagattct    300
gacagttttc atgagaactt gcttttcttt tctctctcaa gtccttattc agtattcagc    360
acccttaaca gattagtccc actgctgagt caggcctctt gcatgaagca gcaatgagaa    420
agacacactt ggccaatgtt atcctggagt aattctcaat gatgccttct ctgtgtttct    480
tcaagacaac tgtccttagt gtgagaaaat gtccagattt ctcacctctt ggccaatgga    540
aggttgtcac tcagagcaac ggcaaggagc agagtgtgtg ctttgacgca gttatggttt    600
gcagtggcca ccacattcta cctcatatcc cactgaagtc atttccagggt gagaccgct    660
gggattccca gctttttgga gtaggtttcc aggtacttta tatgtagttt ggattgacaa    720
gcaggattca ttgctgcaac tgggcagaac ttggctcaat aagattgaga cagagctaga    780
aagatgaaag acaccaaaaca tcatctttgt ttctattggc ctctgagtct tcatcacaca    840
tagatctcag agccaacttc cttggaagtc actaagtcct tggcataatt ttagagaatt    900
cacatcaaac tggttctctg ttggagaggc ctttttagcc atgtgcctgc gttggccttt    960
ttctaccctg ccaaaccacc agcctttttc acagggccat a                          1001

<210> 447
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 2-39-27 : deletion ACAATGACAGCATCATCATCA

<220>
<221> misc_binding
<222> 481..500
<223> 2-39-27.mis1, potential

<220>
<221> primer_bind
<222> 475..495
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 870..892
<223> downstream amplification primer, complement

<400> 447
ccagggacaa gttaagtccc accattagtg taccaaatcc cagtacttac atacttacct    60
tgatgtgtat atgctgcaca ccaaggtaaa ttacaaatca aacatattct gatttttgaa    120
aacataactg aactaatagt cattttaaag gcttactctc tgggtataat taagcttgaa    180
atcatccaaa tattattatt ttcatagctg cagcaaagca gctgaatttc acagagctaa    240
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tgagcaagca gtatactcag tgatacacag gacctccagg tcaaagggtt ataccatgtt    360
gaaaatcaag gaaaaagaca tattcttttc gaaatttctc tctgcttcag tccaccaacc    420
tctcatggaa tttattctct gtacctgtgg gagtcatttc agtgttatga accagcaatc    480
aagacacctt atctgggtcca catcatcac catcatcata tcatcattac tcatcattac    540
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tttttatttt atttatttca atagcttttg ggggagcagg tgggtgtttg ttacctggat    720
aagttcttta atggtgattt gtgggttttg gtgcacccat catctgagca gtgtacacta    780
tacccaatgt gtagtctttt atgcatcacc cccactcca acacttacc ccaagtcccc    840
aaagtccatt gtagtctttt tatgcctttg tattctcata gcctagctcc aacttatgag    900
tgagaacata gaatgttcag tttccactc ccaagttact tcacttagga taatgggtctg    960
caactccatc caggttgctg tgaatgcat tatttcatcc c                          1001

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389

<210> 448  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 2-45-155 : insertion GAC

<220>  
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 <222> 481..500  
 <223> 2-45-155.misl, potential

<220>  
 <221> primer\_bind  
 <222> 347..367  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 764..788  
 <223> downstream amplification primer, complement

<400> 448  
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 ggtagagacc taggatgcta atatcttgca atgtgccaaa ataattgtcc ctgtccccaa 180  
 cctcaccatt gccaatatta cccctacccc tcacagttag cgtcacaggc aggcaacaaa 240  
 ctgggtgtcgt cacagaatga ttgatggaac acatagactg cattcattac ctaaaccattg 300  
 tcgtcacact gcagcaacca aagacaatcg cattaccagc ggggttagatg taggaagagt 360  
 aaaaaacaaa aaatttttga atgcgtaatt atcactaatt attttatttg atccttcagg 420  
 agaattgtgga agatggccga gcaagtatct atcaatctgt cgttaccaac accagcaaag 480  
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 ctaaacttct ggaatatttc aggatttttg ctaaaaaatt tgatctgcta aaatatattc 600  
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 agatagaaaa gttactctga taatgaaagc aattatgaat gaagtatccc attctaagta 720  
 tttgttgaaa tataacagcc tcatataaaa cccaaaaagt agtgtcatta cccttggtat 780  
 tatagattat atacattaat tgaagaggaa aatcatctgt taaaattaaa ggtttgaata 840  
 ataatatatt gatgtcaaaa cttttttttt tttttttctc cctgagacag agtctcactc 900  
 tgttgctcag gctggagtgc agtggcatga tctcagctca ctgcaacctc tgcttccag 960  
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<210> 449  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 2-4-391 : deletion G

<220>  
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 <222> 481..500  
 <223> 2-4-391.misl, potential

<220>  
 <221> primer\_bind  
 <222> 111..132  
 <223> upstream amplification primer

390

<220>  
 <221> primer\_bind  
 <222> 513..537  
 <223> downstream amplification primer, complement

<400> 449  
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 taaagacatt gcattactac tgttgacctc agagcacgcg cctcttgctt aattctagga 180  
 ctccctaacta agtcttttga gtttcagctg gaagaatgct ggaggaatac ggaactcctc 240  
 ccattttctca cagccacctc caactcttaa aaacgcttcc aactgcctcc cagcacacaa 300  
 ccaagggaga aaactattct gtcaaagaga cgggtgccaa aggcaaaaac aaaggtaagg 360  
 atgatcgctg gggaaagaag ctgaaaagga aaagctcaga actctagctg gaaatttggc 420  
 tcacatccct agtatgttac tgcatagtct ggctttgttc aatgggtcgc ttttaaata 480  
 taaagctaga tgtaagcaag gtttgcaaca aagtcataa gaaactcagc ttttctcaa 540  
 ggcaagaaga gagcaggatt tttgactggc tctttattca atagtgtgc ttattaaatt 600  
 accactgcta caatgtttaa agccaattac ctgagcacat cataaggatt ctcttaccgg 660  
 ttgtcccagt taagtaatgt tgattgatca actccttgac aggagctgat ggcaaagaag 720  
 gtagctgtga ttggagctgg ggtcagtgcc ctaatttctc tgaagtgtg tgtggatgag 780  
 ggacttgagc ccacttgctt tgagagaact gaagatattg gaggagtgtg gaggttcaa 840  
 gtaagtgaga ttttcttggg tcttgaacag gttgtgttgt tatttcaggg tgaatcacag 900  
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<210> 450  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 12-345-410 : deletion TAA

<220>  
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 <222> 481..500  
 <223> 12-345-410.misl, potential

<220>  
 <221> primer\_bind  
 <222> 96..113  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 601..621  
 <223> downstream amplification primer, complement

<400> 450  
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 ccctgccaaag ctagcttgac tagatgggtt gtctcataat cctgagacaa cgcagacgct 120  
 ctgcatacct tatcaccacc tttcaacctc ctggggctct gtgattgatt tatttaacga 180  
 cttcagagca gggtgaaaca ctgctctgca tgtctcacta tatatctcac tatttctaga 240  
 tcttcattt ttaaaaggaa ttgggattca ctcatctac aagtattgtt cagcatctac 300  
 tctagagcag atacagtggg gagggaaaaa aagtctcagc ctccatggaa cctataaatc 360  
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 atgaatataa atcctcttaa taaccactac tgcattgcaca aatgaggcct tatacagtat 540  
 gtcgtattgg ggtaaaaaaa aaatcttcta aattaaaaaa acgcaaagtt ggtaggaggg 600  
 cattcaccaa acattaaactg aagtcattct tggcttttaa caccagagta aagtgcattc 660  
 acttttttat tttaaattaa ataccgaagc tgggtgcggg ggctcatgcc tgtaatccca 720

391

```

gcacttggga ggctgaggtg ggcagatcac ttgaggtcag gagttcgaga ccagcctggt 780
caacatggtg aaaccccatc tctactaaaa aaaaaaatac aaaaattagc cgggtatggt 840
ggtgggcgcc tgtaatccca gctacttgag aggggtgaggc aggaaaatca cttgagctca 900
ggagggcgag gttgcagtga gccaaagatca tgccactgca ctccagcctg ggcaacagag 960
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<210> 451  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-358-353 : deletion AGCC

<220>  
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 <222> 481..500  
 <223> 10-358-353.mis1, potential

<220>  
 <221> primer\_bind  
 <222> 149..167  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 577..596  
 <223> downstream amplification primer, complement

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<400> 451
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ttgatttctc tccagaattg tgtttatgct gaccactaaa tatcaactta ttaaaaaaaaa 180
aacttacgtg gttttaattt ttttttcgct cccccctcgc ccacaggaga actaatgaca 240
gttgccagat ggatgagggg gtttatcgca aaccatcctg actacaagca agacagtgtc 300
ataactgatg aaatgaatta tagccttatt ttgaagtgtg accaaattgc aaatgaatta 360
tgtgaatgcc cagagttact tggatcagca tttagggaaag taaaatatag tggaagttaa 420
actgactcat ccaactagac attctacaga aagaaaaatg cattattgac gaactggcta 480
cagtaccatg cctctcagcc agcccggtgtg tataatatga agaccaaagc atagaactgt 540
actgttttct gggccagtga gccagaaatt gattaaggct ttcttttggt ggtaaatcta 600
gagttttatac agtgatcatg tacatagtaa agtatttttg attaacaatg ttttttaata 660
acatatctaa agtcatcatg aactggcttg tacattttta aattccttact ctggagcaac 720
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tacattaaat cacttgaatc cattgaaagt gcttcaaggg taatcttggg tttctagcac 960
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<210> 452  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 10-360-190 : insertion T

<220>  
 <221> misc\_binding  
 <222> 481..500

392

&lt;223&gt; 10-360-190.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 312..329

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 713..731

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 452

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tgattaaggc tttctttggt	aggtaaatct agagtttata	cagtgtacat gtacatagta	120
aagtattttt gattaaca	gtattttaat aacatatcta	aagtcacat gaactggctt	180
gtacattttt aaattcttac	tctggagcaa cctactgtct	aagcagtttt gtaaattgtac	240
tggtaatgtg acaatacttg	cattccagag ttaaaatgtt	tactgtaaat ttttgttctt	300
ttaaagacta cctgggacct	gatttattga aatttttctc	tttaaaaaa ttttctctcg	360
ttaattttcc tttgtcattt	cctttgttgt ctacattaaa	tcacttgaat ccattgaaag	420
tgcttcaagg gtaatcttgg	gtttctagca ccttatctat	gatgtttctt ttgcaattgg	480
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tgatggctgt tatcaaagga	acttttcttt gtttaaattt	gctgatgcag gaattaagtt	600
taaacacaac tctatagaaa	gaaaggagat tattaccag	aattcacatg tagtgattat	660
taaggacaat tttttttttt	aactaaaaaa gttggcggca	ggggtggggg gtggcaatca	720
tttttcttcc tatacatata	aaggatattg tcaaaaatgg	cgttcttctc ttgtggcctg	780
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atagactttc ctctaaaagc	cattcactcc agattttacc	tggggaatat tctacatact	900
gcttactttc tctataaaac	tcatcaataa atcatgaaag	gcaactgagtt ttgtaaatca	960
ggaccctaaa tgtttaattg	taaataagtt tcagataatt	a	1001

&lt;210&gt; 453

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-365-374 : deletion A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-365-374.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 128..145

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 530..548

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 453

tcaaattctt cttttacacg	atgactctta ggtgaccaat	ttatgacgtt tgggtgtctg	60
ttttagtacc ttaacaagaa	atatccgaca taggagagga	gaaaagggtg tcatcaatgt	120
accaagtaag tctactgaga	ggtgggtggg tgggagagag	acatgttgta ttgttggtta	180
atcctggatt ctaaacattt	tttatttttg tatttttata	atacagtatt taaggacaag	240
aatacaccat ctccatttat	agaaacattt actgaggatg	atgaagcttc aagggtcttc	300
aagccggatc atattttacat	ggatgccatg ggatttggaa	tgggcaattg ctgtctccag	360

393

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gtatagtttc aaatatacag agaggcaaag tgttccatcc atttctgttt tttaacttct 420
ttatatatgc atgttttctg ttccaaaaat cacattttta tgaggttgaa atggtagctg 480
gtatgccttt ctgaaaaaca atgaagtat attagtaa atcattggaag ctgtctatga 540
ctaatagttc tacagactct gttgttcacc acaaaggat atacggtata tataccttta 600
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tcaaaactct gaaaatgagc tctgatggaa gtattccaat gattttgtga tccaagggat 780
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gccctgagtg aaagtataca ggagcctgtc tctgtttacc catttaaaaa ttttctgga 900
tctgtttaa tgatgggtcc cctggcagtg agtgtgtcca agctcactta ggaccagagt 960
ggaagagcat ggaacccatc aggtggccac acacaagcct g 1001

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&lt;210&gt; 454

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-367-58 : insertion GTGGAG

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-367-58.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 444..461

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 855..874

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 454

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ttccttctct gacttcactg cttggggagc atgctaata ttccttggtg ttaataactt 60
tgcaattctt aaacaggtga cattccaagc ctgcagtata tctgaggcca gataccttta 120
tgatcagttg gctactatct gtccaattgt tgtaagtaga aattacctct tattttaaat 180
actacttcgt atgaaataag atagcatgtg cagaatttac tgacagtgtg ctatttaagt 240
ccagttaaga cctcagtcag agatggacta atataaatag tatgtaggtt taggtataat 300
gaactgagag tctacactgt agaagtttac tcttgctagt acaacattga tttgttaa at 360
gtgaagtttg aatgtggcca ttttccctcc cccatacttc atgtcctcac attagagaga 420
ggatgattta agtgaatcaa acccaaggaa ctggattctt cctggtatat ttcactgtaa 480
gataggcaca ggtacatgtg ctctgtatgg gttgcataaa catgcttttt gatcagaa at 540
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tttgtttgtt tttagatggc tttgagtgtc gcatctccct tttaccgagg ctatgtgtca 660
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gggaaaaaga tttggtctag gtggggaggt aatttttacc gtatgttgat gtctgtattt 900
gttttagctt catttaaaat tagtagtccg attttctcat attttggaag tgctgatggc 960
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&lt;210&gt; 455

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

394

<222> 501  
 <223> 12-468-424 : insertion T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-468-424.misl, potential

<220>  
 <221> primer\_bind  
 <222> 76..95  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 583..603  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_feature  
 <222> 13,23,255,639,967,971  
 <223> n=a, g, c or t

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 aaacactgta tgtataccag taaatgcttc ctaatggtag atgataaata tttttgttgc 180  
 ttgtagatag aatttttcca tctagtgtaa gcttaggatt gttttctttt tcccagtgac 240  
 atgtacagtt tcacntactc cacttaaaaa aaatcggttag ctcagataaa gtgtgtggca 300  
 catgaaatga attttgccaa ttcaccaccg aaatccctgc tttatatctc ttcagtggaa 360  
 cactaaacca acttctttcc caagtacact gatttgatct ttacaaatta tatgaatgta 420  
 ttaaattatc acatgtgccc tgaaactgtg tacatctatt atgtatcatt agagagatgg 480  
 aaaaaaaaaa aagaaactgc ttagtaaaagt agaaaaaata gttttaaagt aattttgtaa 540  
 ggctaattga agacttactg tataaaacaa aaaggatttt aaccacattc aaattattga 600  
 ctgttttggg gcttttcagg aatcacttaa aaagcaccna agttcacagc caggcacggg 660  
 ggctcatgcc tgtaatccca gcactttggg aggctgaggt gggcagatca cttgaggtca 720  
 ggagtttgag acaagcctgg tcatcatggc aaaatctcat ctctactaaa aatacaaaaa 780  
 ttagacctgg tggcatgtgc ctgtaatccc agctactgag gagtctgagg catgagaaac 840  
 gcttgaacct gggaggcgga ggttgcaagt agccaagatc gtgccactgc attccagcct 900  
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<210> 456  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-481-293 : insertion T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-481-293.misl, potential

<220>  
 <221> primer\_bind  
 <222> 774..793  
 <223> upstream amplification primer, complement

<220>

395

&lt;221&gt; primer\_bind

&lt;222&gt; 333..353

&lt;223&gt; downstream amplification primer

&lt;400&gt; 456

taccattga	actctgatgg	ctgttatcaa	aggaactttt	ctttgtttaa	atttgctgat	60
gcaggaatta	agtttaaaca	caactctata	gaaagaaagg	agattattac	ccagaattca	120
catgtagtga	ttattaagga	caattttttt	ttttaactaa	aaaagttggc	ggcaggggtg	180
gggggtggca	atcatttttc	ttcctataca	tacaaaggat	attgtcaaaa	atggcgttct	240
tctcttggtg	cctgttattc	tgattgctgc	tgtatacagt	tttgctactc	tttagttttt	300
agttaagcat	actgatagac	tttcctctaa	aagccattca	ctccagattt	tacctgggga	360
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agttttgtaa	atcaggaccc	taaatgttta	attgtaaata	agtttcagat	aattattata	480
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attttttcat	cattgcgacc	tcagatgttt	ctagaaagaa	gcatagtgat	gaagacaaac	960
agatgccgta	tctcccctag	tagctatcaa	ccccgggtggg	g		1001

&lt;210&gt; 457

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-499-86 : deletion GGG

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-499-86.mis1, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 567..586

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 136..156

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 77,302,451,665

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 457

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396

```

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```

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<210> 458
<211> 1001
<212> DNA
<213> Homo Sapiens

```

```

<220>
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<222> 501
<223> 12-500-217 : insertion CAATA

```

```

<220>
<221> misc_binding
<222> 481..500
<223> 12-500-217.misl, potential

```

```

<220>
<221> primer_bind
<222> 286..306
<223> upstream amplification primer

```

```

<220>
<221> primer_bind
<222> 714..734
<223> downstream amplification primer, complement

```

```

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<210> 459
<211> 1001
<212> DNA
<213> Homo Sapiens

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<220>
<221> allele
<222> 501
<223> 12-511-101 : deletion A

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397

<220>  
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 <222> 481..500  
 <223> 12-511-101.misl, potential

<220>  
 <221> primer\_bind  
 <222> 585..602  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 152..172  
 <223> downstream amplification primer

<220>  
 <221> misc\_feature  
 <222> 129,588,664  
 <223> n=a, g, c or t

<400> 459  
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<210> 460  
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 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <222> 501  
 <223> 12-586-443 : deletion C

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-586-443.misl, potential

<220>  
 <221> primer\_bind  
 <222> 59..78  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 523..543  
 <223> downstream amplification primer, complement

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<400> 460
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<210> 461
<211> 1001
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-593-287 : insertion TAAAT

<220>
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<222> 481..500
<223> 12-593-287.mis1, potential

<220>
<221> primer_bind
<222> 771..788
<223> upstream amplification primer, complement

<220>
<221> primer_bind
<222> 251..271
<223> downstream amplification primer

<220>
<221> misc_feature
<222> 135,300,327,765,797
<223> n=a, g, c or t

<400> 461
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399

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&lt;210&gt; 462

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-795-383 : insertion CA

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-795-383.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 119..136

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 679..696

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 408,416..417

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 462

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&lt;210&gt; 463

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

400

&lt;223&gt; 10-494-332 : insertion ATAAAA

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-494-332.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 170..188

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 574..591

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 463

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&lt;210&gt; 464

&lt;211&gt; 1001

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-494-380 : deletion GATAAT

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-494-380.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 122..140

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 526..543

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 464

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401

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&lt;210&gt; 465

&lt;211&gt; 929

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 429

&lt;223&gt; 12-659-251 : deletion TT

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 409..428

&lt;223&gt; 12-659-251.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 179..198

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 611..631

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 80,605

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 465

```

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402

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929

&lt;210&gt; 466

&lt;211&gt; 987

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-912-419 : deletion A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-912-419.mis1, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 83..103

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 554..574

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 86

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 466

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gtctctgaga catttgttgt atcaaagcac aacaacatta acagtaaaac aaaaaatatt	660
gtgaaatatg aaatatctga tcctttctcc ctctctcaaa aattctgtct gccagggtgtg	720
gtggctcacg cctgtaatcc cagcactttg ggatgacaag atgtttggat cacctgaggt	780
caggagttag agaccagact ggacaatgag gtaaaactcc gtccctgcta aaaatacaaa	840
aattaaccag gcatggtagc acatgcctgc aatcccagct actcaggaga ctaaggcaca	900
agaattgcat gaaccaagaa ggcagaagtt gcagtgaagt gagatcgag cactgcactc	960
tggcctgggc aacagagcaa gactccg	987

&lt;210&gt; 467

&lt;211&gt; 806

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 306

&lt;223&gt; 12-914-28 : deletion T

&lt;220&gt;

&lt;221&gt; misc\_binding

403

<222> 286..305  
 <223> 12-914-28.misl, potential

<220>  
 <221> primer\_bind  
 <222> 279..298  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 773..793  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_feature  
 <222> 536  
 <223> n=a, g, c or t

<400> 467  
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 atcttttagga ttgctgattc cttactctgt ttgaccaagt ctgctgttga agttttctac 180  
 tgagtttttc aattcacttc acatatctttt tattaattgg atttcttttt attttttaaaa 240  
 catttccact tctttgcaaa atttatcatc attttcctgg attattctct aaagtgtggt 300  
 ttttttcaac tctctattca tatgtgctgt gatttctgaa cttttaaaat aggtgtgcac 360  
 tgaatttctt gtcaaaaatt ttatggagca tctgaacctt aattctgcac tgtaacttaa 420  
 aactagactt gcagtgattt ctaggtctgt gagatactta agcaataact ggagcttaat 480  
 ctcaaatggt atacttggtt gcaggtcata gaaaagctcc gtatgagtaa ctgggnatta 540  
 taaaaatacc tgccaaagat tcaggccttc ctttggtatt tggcccctgc agtggtatgg 600  
 aactggtcag actcctcagc gtggcattcc tgctaataagg aacaccaagc agttgctgag 660  
 atgtgtgtgc ctgtcactgt gattagtact tccactcttg tttccaattc accccaggtc 720  
 attcaactct atagccactc ctaatgcctc tcttgaggta ggacaaaagt gggcttccca 780  
 aaccaaaaag attcacagat ccatga 806

<210> 468  
 <211> 1001  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-624-307 : deletion T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-624-307.misl, potential

<220>  
 <221> primer\_bind  
 <222> 788..808  
 <223> upstream amplification primer, complement

<220>  
 <221> primer\_bind  
 <222> 340..360  
 <223> downstream amplification primer

<220>  
 <221> misc\_feature  
 <222> 132,690,865  
 <223> n=a, g, c or t

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<400> 468
tgagatagca tgttcttggg ttttaatata atattatcaa tccattagtg ccatttagaa      60
aacttaaatt gaccagcaat ccggtgctct ctatctgata aatttacagc ctgtttttgg      120
acacctactt gnataatttta aatatttatt aataaatata ttagaatgtt aattaaaatt      180
tttaattatg gctaaaattt tagtaaacaa atttttaaaa ttttaatgtc ttatgtcaac      240
attttatgaa ttctacagta gtttagtgat ttcttcta attttaagggtt atatttgtag      300
taattttgag atgtgattac aaatgcattg gcaatcctct catcaagtca tatacccttc      360
ctgtgaaatt tagcaaattt tgtggctgtc tcaaaaaaga gaatatagtg gaagttagga      420
gcttaacttc caaagctcaa aaaaattgtt aatacatttt ctctgtctgt ccttctgtcc      480
ttttctctgt gtttctgtca ttcttttttc ctttttctct gtctctccac tcgctttgga      540
gccctgtgct atatccacag tccccaggag aaatatacca tttggtgtag aaatacatat      600
gtatgcctaa taagtcacaa ttggccagc actgcagtag cttgagactt ttaatactaa      660
tcaccagata tgtgagtgat taaccttc an gatgattagc ccaagctacc ttaggacagt      720
aaccaactga gaaatgctgt gccataaaca cctaattgag cctggccaac aacctagaat      780
taaaacagat gatgataaaa tgttgctgtt ctaagcacct tcatttcaga gcagttgggt      840
aagcagtgat agacaaccag aaaaatattt ttacatattt tgagttatac tcctgtaata      900
atattaaaat taatatatac ctacaaaaaa agtgaatcta tattacattt tgtaactacc      960
tgtaaccag aaagagtagt cagcatagga tgggtgggtaa g                                1001

<210> 469
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-290-326 : polymorphic base A or G

<220>
<221> misc_binding
<222> 481..500
<223> 10-290-326.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-290-326.mis2, potential complement

<220>
<221> primer_bind
<222> 177..196
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 576..595
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-290-326 potential probe

<400> 469
atgctttgaa tgttcaaaga aaccaagtga tctctatggc taggcattcc ctttaactta      60
tgtgatgagc taaaattttc aaattgcaaa aatatagaag ttctggacag tgtaacgat      120
aggacagcct acctcaggac tctcagcttc cttattacag tccagtttgt cactttttctc      180
cctcatgtcc actctgcagg atcaccggtt tccgactgag tctggggatt ttggccttgt      240
tgacctcctc aggtgccctg ggaattgcaa acagctttct ggatgaatat ctggacctca      300
atattgcaa gaaactgagg cggcaattct aactttttct cttcccttta atgcttgtag      360
aagctgttcc caccatgaag gtaatatggt atcatttgtt aaataaaaat aaagtcttta      420

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405

ttctgttttt	cttgaaatgg	ctttgtagaa	acacacactt	tagagaatac	atttcctgtt	480
taagaactga	aatttgcttg	rggtaggtac	atccatatca	tcagtgggaag	tgtgttggtgc	540
cctagtttta	gaaatgcatg	ctgaagtcac	gaccggttgt	gtagggacac	agcagcctga	600
cccgtgtgct	ggccgttccg	aggactactg	acatatcctg	ccactctgac	tgttctctc	660
ttgatggaaa	gtttcagaca	gaagctgata	aagcagctga	gccctttcat	tggttatagg	720
aaaacaacag	ataggaaggc	atgtggggga	catcaggcaa	acacatagag	tagccagaga	780
agggagtcac	caactgatag	cacaaaacaa	aacagggttat	agaaggctgg	gcgtgggtggc	840
tcatgcctgt	aatcccagca	ctttgagaag	ccgaggcggg	cagatcacia	ggtcaggaga	900
tcgagaccat	cctggctaac	acagtgaac	cctgtctcta	ctaaagatac	aaaaaattag	960
ccgggcatgg	tggcgggtgc	ctgtagtcc	agctactcgg			1000

&lt;210&gt; 470

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-290-37 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-290-37.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-290-37.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 465..484

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 864..883

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-290-37 potential probe

&lt;400&gt; 470

aaaaaacggt	aaggagaacc	cagtaatttt	gtattttatgc	aaaaagtagc	aggataaggt	60
ctgggtcagt	ttccttctcc	tgagagatat	taacagcact	gtgagttgtg	gtggcaaaaag	120
taatccatac	tgtctgaggt	caagtcttgc	cagttcatac	acaattaggc	atgagatggg	180
aaaatgcaag	tataaaactg	tgactgaat	gtggactcta	gttccttgca	aaaatttttgc	240
attgtctcca	gaaagttacc	gaggaaataa	tcttgccctag	ccttgataat	gctttgaatg	300
ttcaaaagaa	ccaagtgatc	tctatggcta	ggcattccct	ttaacttatg	tgatgagcta	360
aaattttcaa	attgcaaaaa	tatagaagtt	ctggacagtg	ttaacgatag	gacagcctac	420
ctcaggactc	tcagcttcc	tattacagtc	cagtttgtca	cttttctccc	tcattgtccac	480
tctgcaggat	caccggtttc	ygactgagtc	tggggatttt	ggccttggtg	accctcctag	540
gtgccctggg	aattgcaaac	agctttctgg	atgaatatct	ggacctcaat	attgccaaga	600
aactgaggcg	gcaattctaa	cttttctct	tccctttaat	gcttgacagaa	gctgttccca	660
ccatgaaggt	aatatgggat	catttggtta	ataaaaaata	agtctttatt	ctgtttttct	720
tgaaatggct	ttgtagaaac	acacacttta	gagaatacat	ttcctgttta	agaactgaaa	780
tttgcttggg	gtaggatcat	ccatatcatc	agtggagtg	tggttgcccc	tagttttaga	840
aatgcatgct	gaagtcatga	ccggttggtg	agggacacag	cagcctgacc	cgtgtgctgg	900
ccgttccgag	gactactgac	atatcctgcc	actctgactg	cttctctctt	gatggaaagt	960

406

ttcagacaga agctgataaa gcagctgagc ccttttcattg

1000

&lt;210&gt; 471

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-523-232 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-523-232.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-523-232.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 270..288

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 527..545

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-523-232 potential probe

&lt;400&gt; 471

acttcaggag	attttaaggg	aattatttaa	agaccatttt	gtaatgttct	gggtaactca	60
agtggaaagg	aatggatgtg	agggattttt	taaaataaga	aatctcttga	atatgaattg	120
agtaatctct	taagttcttt	agataagata	tggagtgatc	tttaggctca	ggaggcctgt	180
cttttgaggaa	agtgtctataa	ctcacttcta	ctaccggcaa	cttaatacgt	aagagaatat	240
tcttgccct	gctctgaggt	ttaaacaatg	ttcaaccttg	aaagattgat	tttaaattga	300
aaggtcattt	aaactctttt	gtaagttctt	atattttgtt	aaaataggct	gaacattttac	360
ctctagatga	aaaatcaatg	cttcagtgtg	tacttttctc	ataaaataac	ctatgtcttt	420
atttttttct	gcagacaaaa	ctgtgtggag	ttttatccta	tattcataat	tacattgttg	480
atggctgggt	ggtatttcaa	ycaggtaat	gttaaaatac	agtcaactgt	gttctaattga	540
tgactatgat	gcttaaacga	ttaaggagta	gatatatggc	caggcacggt	ggctcacacc	600
tatattccca	gcactttggg	aggcccaggt	gggtggatca	cctgagggtca	ggagttcgag	660
accagcctgg	ccaaaatggg	gaaaccctgt	ctctactaaa	aatacaaaaa	ttagctaagt	720
gtggtggcgc	atgcctgtaa	tcccagctac	ttgggaggct	gagacaggag	aatcacttga	780
acccaggagg	tggaggctgc	agtgaactga	gatcgtgcca	ctacactcca	gtctgggtga	840
cagagactcc	atctcaaaaa	aacaacaaaa	aaaaggagca	gatatgcaac	tatatagcta	900
ccaatcctgg	agactaaaca	aaataaacag	tgtgtgcatc	agagtgtctat	gttgcaggta	960
tatgaacttt	ggcttcattc	taatttaatt	caataatgaa			1000

&lt;210&gt; 472

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

407

<222> 501  
<223> 12-449-300 : polymorphic base T or C

<220>  
<221> misc\_binding  
<222> 481..500  
<223> 12-449-300.mis1, potential

<220>  
<221> misc\_binding  
<222> 502..521  
<223> 12-449-300.mis2, potential complement

<220>  
<221> primer\_bind  
<222> 783..800  
<223> upstream amplification primer, complement

<220>  
<221> primer\_bind  
<222> 325..345  
<223> downstream amplification primer

<220>  
<221> misc\_binding  
<222> 489..513  
<223> 12-449-300 potential probe

<400> 472  
ccaccagagt tcataccagc atggaatctg tacacatctt taaatgatac ttagatttac 60  
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actaactctt tctccaaaag aggatccaaa gtccctttat ccctctttgg gccatgatta 180  
tttgtcttct aacagagtca cattgtctcc ttgagacgta aagatgcatt tagatgcagg 240  
gggacggaag aaaggaaaag aaaaaatagt ttatatgtat atacatttgt atcaaaataa 300  
gaaataaata ctcataatta ttacaatttt cctctgaaaag tagtcacatg gttataacctg 360  
gtattttattt attttttattt attattatta ttatttgaga cagggtctca ccctgtcaca 420  
caggctggag tgcagtgggtg caattatggc tctctgctac ctctgccacc tgggctcaag 480  
ccatcctctc accttagcct yctgaacagc tgggatacat gtgcactaat tttttttttt 540  
ttgaaacaaa aatatattgtg tagaaggcac aaaagctaca atcacagact ccactgtgca 600  
aaggcgcaac ctgcctcatt gatctctagt gtacaccaac ccagcttccc tttccattca 660  
gcctgtgaaa ggagatagtg cttgggccat ttggtaaaag aaggggatgg gagatgatca 720  
aaaccccaag taaggttcat ccaatatggg gtctaagcag caaatgacta attgctgaag 780  
aaggagacta gacagaggat tagaggcagc catggggcca gtgcagctgt ggagagctct 840  
gagcaaagaa acaagggttg caggtgagga ggcctaggat agaggccaga aggccaaagcc 900  
tggggctgca tgtgcaactaa tttttgtatt tttttgcttt tttgatagag atgagatttc 960  
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<210> 473  
<211> 432  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> allele  
<222> 212  
<223> 10-186-212 : polymorphic base G or C

<220>  
<221> misc\_binding  
<222> 192..211  
<223> 10-186-212.mis1, potential

<220>

408

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<221> misc_binding
<222> 213..232
<223> 10-186-212.mis2, potential complement

<220>
<221> primer_bind
<222> 1..20
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 413..432
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 200..224
<223> 10-186-212 potential probe

<400> 473
agtatgacga taggcagatg ttgtcttgca aaacgagcaa acaggcatat gaaatcagat      60
acctagctgc caaactatat gtgaggcatt tccgaagaat ttaacgacct cgatagagcg      120
cagtcaagtt tggatgaacag aatatgtctc tgaactagag gaggcctcac acaaggagta      180
gggtcagacc ccgagtgga ggaggagga gsagtagaaa cagtccagct cgccgcccac      240
gtaacctggg tcctgaatcg gcccgcttg gccagtgtc cagaagcgcg gagaggaaacg      300
ggctggggcc caaaaaagag gggggagcct gaacgtccgg gggaaagttc ggaggcgcg      360
gaaccacgg atggaacct gtctttgggg aaaaggacca cacctgtcag cagagtcctg      420
cagacgtgag aa                                     432

<210> 474
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-286-289 : polymorphic base G or C

<220>
<221> misc_binding
<222> 481..500
<223> 10-286-289.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-286-289.mis2, potential complement

<220>
<221> primer_bind
<222> 213..231
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 613..631
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-286-289 potential probe

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<400> 474
aaaaaaaaa gtatgattat tatcagagtc tctaggtgac agtgagaggg agcctgggtgc      60
aggggatagg gcacagttgt aatctgctag actgggttca aatcctaaat cggccattca      120
gagcgtgaat acattagaac tggcttaaata tagcatttaa aatgtcaaca ggattaagta      180
agtagttcct tctcactttt tccacaggag gtgagaaaagg tctcagcaag gccctaggag      240
ggaagggcgt ggggatgaaa gggatccaga gagtctgtgc attggcagag aagatagtgt      300
aactggcact gcggctggag ggctgggtccc acagtggagct acagccctgc ccgctggcgc      360
tgggagagggc ttaaaacaaa cgccggaagc aactcccagc ccataaaaga tctgtgaccg      420
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aaagatggcc ggggaactcga tcctgctggc tgctgtctct attctctcgg cctgtcagca      660
aagtaagagg catgggaagt tcgtgtgtgt gcgctgtgtg gcgtgtgtgt gtgtgtgtga      720
caaggcttgc gggagagaga gggagggagg gagatgggtc cgggtgtttg ttctctactt      780
gcccttgtag gtagctctgg gtccctcagag cacagtcgcc tcagggtcac ccatgccgcc      840
tgctaccctc cttcccaggg gcaagcagag actgagaaca ttccagagat tagttctccc      900
aactggaacg ctgtggggcc tcagagctca gcgattctgc atcatctgtg attacgacct      960
acagcccggt caaacgagcg ttagtagcct gtaacctgc      1000

<210> 475
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 10-286-345 : polymorphic base A or T

<220>
<221> misc_binding
<222> 481..500
<223> 10-286-345.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 10-286-345.mis2, potential complement

<220>
<221> primer_bind
<222> 158..176
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 558..576
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 10-286-345 potential probe

<400> 475
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attcagagcg tgaatacatt agaactggct taaattagca tttaaaatgt caacaggatt      120
aagtaagtag ttccttctca ctttttccac aggaggtgag aaagggtctca gcaaggccct      180
aggaggggaag ggcgtgggga tgaaagggat ccagagagtc tgtgcattgg cagagaagat      240
agtgtaactg gcaactgcggc tggagggctg gtcccacagt gagctacagc cctgcccgc      300
ggcgtggga gaggtttaa acaaacgccg gaagcaactc ccagcccat aaagatctgt      360
gaccggcagc cccagacctg cctgccttcc tgacttctgt tccagagcaa aggtcattca      420

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410

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gccgcttgaa tcagcctttt ccccccaccc ggtccccaac tttgtttacc cgataaggaa 480
ggtcagcatt caaagtcaag wagcgccatt tatcttcccc tgcgctctac aaatagttcc 540
gtgagaaaga tggccgggaa ctcgatcctg ctggctgctg tctctattct ctcggcctgt 600
cagcaaagta agaggcatgg gaagttcgtg tgtgtgcgcg tgtgtgcgtg tgtgtgtgtg 660
tgtgacaagg cttgcgggag agagagggag ggagggagat gggtcgggtg ttttgtttcc 720
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ccgcctgcta ccctccttcc caggggcaag cagagactga gaacattcca gagattagtt 840
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gaccacagc ccgttcaaac gagcgtagt agcctgctaa cctgcaggaa gtggtgtgaa 960
tattaattac aagtgttcca aaggaaacgt gcctgcttct 1000

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&lt;210&gt; 476

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-286-375 : polymorphic base A or G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-286-375.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-286-375.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 128..146

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 528..546

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-286-375 potential probe

&lt;400&gt; 476

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gctagactgg gttcaaatcc taaatcggcc attcagagcg tgaatacatt agaactggct 60
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411

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1000

&lt;210&gt; 477

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 10-289-201 : polymorphic base C or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 10-289-201.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 10-289-201.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 307..324

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 700..719

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 10-289-201 potential probe

&lt;400&gt; 477

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&lt;210&gt; 478

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

412

<222> 501  
 <223> 10-321-28 : polymorphic base A or G  
  
 <220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 10-321-28.mis1, potential  
  
 <220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 10-321-28.mis2, potential complement  
  
 <220>  
 <221> primer\_bind  
 <222> 474..491  
 <223> upstream amplification primer  
  
 <220>  
 <221> primer\_bind  
 <222> 890..909  
 <223> downstream amplification primer, complement  
  
 <220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 10-321-28 potential probe

<400> 478  
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 gctgctgctg ggggtggcccc gacccaagat ttacctggga atgggtagcc tcactcagaa 180  
 ggggtgcccctg atgtgggggc acaggtgggt ctttggggac ccctcctggg tgggtgccagg 240  
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<210> 479  
 <211> 422  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 265  
 <223> 10-98-265 : polymorphic base A or G

<220>  
 <221> misc\_binding  
 <222> 245..264  
 <223> 10-98-265.mis1, potential

<220>



413

<221> misc\_binding  
 <222> 266..285  
 <223> 10-98-265.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 1..19  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 404..422  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 253..277  
 <223> 10-98-265 potential probe

<400> 479  
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 tacacggttc aggatttgac acagtcacca cagccatctc ctggagcctc atgtaccttg 180  
 tgaccaagcc tgagatacag aggaagatcc agaaggwgct gggtacatgg gggcccccaa 240  
 ccctatagcc aggagaagcc ttgaraccca ggttggttgt tcagtctaca aacacctgtt 300  
 atgtgcctgc tgtgtgcaag ccctgggcac acagtagtgc ctgcccttgc ctagaagatg 360  
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 ta 422

<210> 480  
 <211> 1000  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> allele  
 <222> 501  
 <223> 12-157-115 : polymorphic base C or T

<220>  
 <221> misc\_binding  
 <222> 481..500  
 <223> 12-157-115.mis1, potential

<220>  
 <221> misc\_binding  
 <222> 502..521  
 <223> 12-157-115.mis2, potential complement

<220>  
 <221> primer\_bind  
 <222> 387..407  
 <223> upstream amplification primer

<220>  
 <221> primer\_bind  
 <222> 833..853  
 <223> downstream amplification primer, complement

<220>  
 <221> misc\_binding  
 <222> 489..513  
 <223> 12-157-115 potential probe

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<400> 480
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aggccctctc gcacctgagg tgagctgggt tccccaccctc accccatccc aaggggtagg      180
gaaaagtcca agaccattcc tcggtgtctcc ttcagggggt gtgcctcctt tttcctctcc      240
tccctctgta gcagggggtct cacttgccgg ctccccaggg caatgggcca gcctagatga      300
tgggggtcgg cctcagccac tcttttctat aattctcagc cagatttcaa gttgggtgct      360
tctccttgga tttatggcac atgggtgggc tgcctttcca ttgattgggg atattttgag      420
caaaataaca gtgataacac tgaagagcat tgcaagcatc cttcaagatg agtcaggctc      480
agtgagttag ttgaaggggt yagtcacta tttgttgata ccactggtac aaaatgatct      540
aaagcccaag acttaccttc ctggagatgc caatccattg gtggggaggg ggaaagggga      600
agaagttagt gatattataa attagtattt aaagagtttg ggtcatctct aagaatacct      660
ccccatatg ggaagattca ttacttaggc gcatagcaga actatctgtt actttaagct      720
gcatacaaca ggcagatttt aaaataaatg tcatgaacat caactaagca caatgatag      780
ctgagtactg aagggacaca gagatacata agacctagtc ccagcctttt aaggacttag      840
agataagaca gacacataca aacaggatta taggtcaaag ctgactaaca ggactgtgtt      900
tacaggtaaa cccacacaca catcagcaaa aaaatatctg agaacaaaaa aaaaggggct      960
actaacaaaa gcaactcagt ttatacaaaa ggagctatca                                1000

<210> 481
<211> 1000
<212> DNA
<213> Homo Sapiens

<220>
<221> allele
<222> 501
<223> 12-472-48 : polymorphic base G or T

<220>
<221> misc_binding
<222> 481..500
<223> 12-472-48.mis1, potential

<220>
<221> misc_binding
<222> 502..521
<223> 12-472-48.mis2, potential complement

<220>
<221> primer_bind
<222> 454..472
<223> upstream amplification primer

<220>
<221> primer_bind
<222> 919..939
<223> downstream amplification primer, complement

<220>
<221> misc_binding
<222> 489..513
<223> 12-472-48 potential probe

<400> 481
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tggggagcat gctaattgatt ccttgggtgt aataactttg caattcttaa acaggtgaca      120
ttccaagcct gcagtatata tgaggccaga tacctttatg atcagttggc tactatctgt      180
ccaattgttg taagtagaaa ttacctctta ttttaaatac tacttcgtat gaaataagat      240
agcatgtgca gaatttactg acagtgtgct atttaagtc agttaagacc tcagtcagag      300
atggactaat ataaatagta tgtaggttta ggtataatga actgagagtc tacactgtag      360
aagtttactc ttgctagtac aacattgatt tgtaaagtgt gaagtttgaa tgtggccatt      420

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415

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ctgtatgggt	tgataaaca	tgcttttga	tcagaaatat	aactgcatgg	agcttttttt	600
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tgatttctgc	atctgtagat	gatagaactc	gggaggagcg	aggactggag	gtgggaattg	780
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ggggagttaa	tttttatcgt	atgttgatgt	ctgtatttgt	tttagcttca	tttaaaatta	960
gtagtccgat	ttttctatat	tttggaagtg	ctgatggctg			1000

&lt;210&gt; 482

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-477-151 : polymorphic base G or A

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-477-151.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-477-151.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 631..651

&lt;223&gt; upstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 113..133

&lt;223&gt; downstream amplification primer

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-477-151 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 419

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 482

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ctctgataat	aagtatatcc	attatctcca	aaaatgtcat	catgcccttt	tggttaatttc	600
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416

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&lt;210&gt; 483

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 501

&lt;223&gt; 12-479-214 : polymorphic base G or T

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..500

&lt;223&gt; 12-479-214.mis1, potential

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 502..521

&lt;223&gt; 12-479-214.mis2, potential complement

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 288..305

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 753..773

&lt;223&gt; downstream amplification primer, complement

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 489..513

&lt;223&gt; 12-479-214 potential probe

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 519

&lt;223&gt; n=a, g, c or t

&lt;400&gt; 483

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tcttagatct gcctcctctc ctgttactcc cccttctctc actccactcc agccatacta	120
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417

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&lt;210&gt; 484

&lt;211&gt; 1000

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 500

&lt;223&gt; 10-290-328 : deletion of G

&lt;220&gt;

&lt;221&gt; misc\_binding

&lt;222&gt; 481..499

&lt;223&gt; 10-290-328.misl, potential

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 174..193

&lt;223&gt; upstream amplification primer

&lt;220&gt;

&lt;221&gt; primer\_bind

&lt;222&gt; 573..592

&lt;223&gt; downstream amplification primer, complement

&lt;400&gt; 484

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gatgagctaa aattttcaaa ttgcaaaaat atagaagttc tggacagtgt taacgatagg 120  
acagcctacc tcaggactct cagcttcctt attacagtcc agtttgtcac ttttctccct 180  
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acaacagata ggaaggcatg tgggggacat caggcaaaaca catagagtag ccagagaagg 780  
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&lt;210&gt; 485

&lt;211&gt; 49312

&lt;212&gt; DNA

&lt;213&gt; Homo sapiens

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; 5466..7466

&lt;223&gt; 5'regulatory region

&lt;220&gt;

&lt;221&gt; exon

&lt;222&gt; 7467..7725

&lt;223&gt; exon 1

&lt;220&gt;

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<222> 45728..45965  
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<222> 45966..49312  
<223> 3'regulatory region

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<222> 7564  
<223> 10-286-289 : polymorphic base G or C

<220>  
<221> allele  
<222> 7619  
<223> 10-286-345 : polymorphic base A or T

<220>  
<221> allele  
<222> 7649  
<223> 10-286-375 : polymorphic base A or G

<220>  
<221> allele  
<222> 17258  
<223> 12-425-57 : polymorphic base A or G

<220>  
<221> allele  
<222> 21590  
<223> 12-421-135 : insertion of T

<220>  
<221> allele  
<222> 21595  
<223> 12-421-140 : polymorphic base A or G

<220>  
<221> allele  
<222> 36971  
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435

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&lt;210&gt; 486

&lt;211&gt; 750

&lt;212&gt; DNA

&lt;213&gt; Homo sapiens

&lt;220&gt;

&lt;221&gt; 5'UTR

&lt;222&gt; 1..201

&lt;220&gt;

&lt;221&gt; CDS

&lt;222&gt; 202..645

&lt;220&gt;

&lt;221&gt; 3'UTR

&lt;222&gt; 646..750

&lt;220&gt;

&lt;221&gt; polyA\_signal

&lt;222&gt; 721..726

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 98

&lt;223&gt; 10-286-289 : polymorphic base G or C

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 153

&lt;223&gt; 10-286-345 : polymorphic base A or T

&lt;220&gt;

&lt;221&gt; allele

&lt;222&gt; 183

&lt;223&gt; 10-286-375 : polymorphic base A or G



436

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<222> 426
<223> 10-523-232 : polymorphic base C or T
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<222> 478
<223> 10-289-201 : polymorphic base C or T
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<223> 10-290-37 : polymorphic base C or T
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<223> n=a, g, c or t
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cccgataagg	aaggtcagca ttcaaagtca agwagcgcca tttatcttcc cgtgcgctct	180
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Val Ser Ile Leu Ser Ala Cys Gln Gln Ser Tyr Phe Ala Leu Gln Val		
	15 20 25	
gga aag gca aga tta aaa tac aaa gtt acg ccc cca gca gtc act ggg		327
Gly Lys Ala Arg Leu Lys Tyr Lys Val Thr Pro Pro Ala Val Thr Gly		
	30 35 40	
tca cca gag ttt gag aga gta ttt cggt gca caa caa aac tgt gtg gag		375
Ser Pro Glu Phe Glu Arg Val Phe Arg Ala Gln Gln Asn Cys Val Glu		
	45 50 55	
ttt tat cct ata ttc ata att aca ttg tgg atg gct ggg tgg tat ttc		423
Phe Tyr Pro Ile Phe Ile Ile Thr Leu Trp Met Ala Gly Trp Tyr Phe		
	60 65 70	
aay caa gtt ttt gct act tgt ctg ggt ctg gtg tac ata tat ggc cgt		471
Asn Gln Val Phe Ala Thr Cys Leu Gly Leu Val Tyr Ile Tyr Gly Arg		
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His Leu Xaa Phe Trp Gly Tyr Ser Glu Ala Ala Lys Lys Arg Ile Thr		
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Gly Phe Xaa Leu Ser Leu Gly Ile Leu Ala Leu Leu Thr Leu Leu Gly		
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gcc ctg gga att gca aac agc ttt ctg gat gaa tat ctg gac ctc aat		615
Ala Leu Gly Ile Ala Asn Ser Phe Leu Asp Glu Tyr Leu Asp Leu Asn		
	125 130 135	
att gcc aag aaa ctg agg cgg caa ttc taa ctttttctct tccctttaat		665
Ile Ala Lys Lys Leu Arg Arg Gln Phe *		
	140 145	
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<212> DNA
<213> Homo sapiens
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<220>



437

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acraatagtt ccgtgagaaa g atg gcc ggg aac tcg atc ctg ctg gct gct      231
                               Met Ala Gly Asn Ser Ile Leu Leu Ala Ala
                               1           5           10
gtc tct att ctc tcg gcc tgt cag caa aac aaa act gtg tgg agt ttt      279
Val Ser Ile Leu Ser Ala Cys Gln Gln Asn Lys Thr Val Trp Ser Phe
                               15           20           25
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Ile Leu Tyr Ser *
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```

438

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 <211> 147  
 <212> PRT  
 <213> Homo sapiens

<220>  
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 <222> 93  
 <223> Xaa=Tyr or His

<220>  
 <221> VARIANT  
 <222> 109  
 <223> Xaa=Arg or Stop

<400> 488

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20          25          30
Tyr Lys Val Thr Pro Pro Ala Val Thr Gly Ser Pro Glu Phe Glu Arg
35          40          45
Val Phe Arg Ala Gln Gln Asn Cys Val Glu Phe Tyr Pro Ile Phe Ile
50          55          60
Ile Thr Leu Trp Met Ala Gly Trp Tyr Phe Asn Gln Val Phe Ala Thr
65          70          75          80
Cys Leu Gly Leu Val Tyr Ile Tyr Gly Arg His Leu Xaa Phe Trp Gly
85          90          95
Tyr Ser Glu Ala Ala Lys Lys Arg Ile Thr Gly Phe Xaa Leu Ser Leu
100         105         110
Gly Ile Leu Ala Leu Leu Thr Leu Leu Gly Ala Leu Gly Ile Ala Asn
115         120         125
Ser Phe Leu Asp Glu Tyr Leu Asp Leu Asn Ile Ala Lys Lys Leu Arg
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Arg Gln Phe
145

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<210> 489  
 <211> 30  
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<400> 489

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Met Ala Gly Asn Ser Ile Leu Leu Ala Ala Val Ser Ile Leu Ser Ala
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Cys Gln Gln Asn Lys Thr Val Trp Ser Phe Ile Leu Tyr Ser
20          25          30

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 <213> Homo sapiens

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439

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&lt;210&gt; 491

&lt;211&gt; 274

&lt;212&gt; DNA

&lt;213&gt; Homo sapiens

&lt;400&gt; 491

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&lt;210&gt; 492

&lt;211&gt; 18

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; sequencing oligonucleotide PrimerPU

&lt;400&gt; 492

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18

&lt;210&gt; 493

&lt;211&gt; 18

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; sequencing oligonucleotide PrimerRP

&lt;400&gt; 493

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18